	FILE 'REGISTRY' ENTERED AT 11:27:33 ON 07 OCT 2010
	EXP HYDROXYETHYL/CN
	EXP HYDROXYETHYLSTARCH/CN
	EXP HYDROXYETHYL STARCH/CN
T.1	1 S E3
шт	EXP HYDROXYETHYL AMYLOPECTIN/CN
т О	
L2	1 S E3
L3	1 S E4
	FILE 'HCAPLUS' ENTERED AT 11:28:39 ON 07 OCT 2010
L4	2856 S L1-L3
L5	199427 S (STERILE OR STERILIZATION OR PHYSIOLOGICAL OR INTRAVENOUS)
L6	170 S L4 AND L5
L7	406196 S (MOLECULAR WEIGHT OR MW OR DALTON OR KILODALTON OR KDA OR DA)
L8	10 S L6 AND L7
L9	1085 S L1/THU OR L2/THU OR L3/THU
L10	102 S L7 AND L9
110	102 6 17 140 13
	FILE 'STNGUIDE' ENTERED AT 12:02:47 ON 07 OCT 2010
	FILE 'HCAPLUS' ENTERED AT 12:03:24 ON 07 OCT 2010
L11	69 S L10 AND (PY<2006 OR AY<2006 OR PRY<2006)
L12	1130548 S INTRAVENOUS OR PLASMA OR (VOLUME EXPANDER) OR (DEGREE OF SUBS
L13	42 S L11 AND L12
	10 0 111 1110 110

=> file reg
COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.22 0.22

FULL ESTIMATED COST

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STRUCTURE FILE UPDATES: 6 OCT 2010 HIGHEST RN 1245698-26-3 DICTIONARY FILE UPDATES: 6 OCT 2010 HIGHEST RN 1245698-26-3

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http://www.cas.org/support/stngen/stndoc/properties.html

```
=> exp hydroxyethyl/cn
            1 HYDROXYETHOXYDIMETHYLSILANE/CN
E1
E2
            1
                 HYDROXYETHOXYETHYL HYDROGEN MALEATE/CN
            0 --> HYDROXYETHYL/CN
Е3
E4
            1 HYDROXYETHYL 1-METHYLALLYL CELLULOSE/CN
E5
            1
                 HYDROXYETHYL 2-(2-HYDROXY-3-(OCTADECYLOXY)PROPOXY)ETHYL 2-(2
                  -HYDROXY-3-SULFOPROPOXY)ETHYL CELLULOSE SODIUM SALT/CN
Ε6
            1
                 HYDROXYETHYL 2-(2-HYDROXY-3-(OCTADECYLOXY)PROPOXY)ETHYL CELL
                  ULOSE/CN
E7
            1
                 HYDROXYETHYL 2-(2-HYDROXY-3-SULFOPROPOXY)ETHYL CELLULOSE SOD
                  IUM SALT/CN
E8
            1
                 HYDROXYETHYL 2-(P-2-PYRIDYLPHENYL) PROPIONATE/CN
E9
            1
                 HYDROXYETHYL 3-(3,5-DI-TERT-BUTYL-4-HYDROXYPHENYL)PROPIONATE
                  /CN
E10
            1
                 HYDROXYETHYL 3-DODECYLOXY-2-HYDROXYPROPYL CELLULOSE PHOSPHAT
                  E/CN
                 HYDROXYETHYL 3-STEARYLOXY-2-HYDROXYPROPYL CELLULOSE/CN
E11
            1
E12
                 HYDROXYETHYL 3-STEARYLOXY-2-HYDROXYPROPYL CELLULOSE PHOSPHAT
            1
                  E/CN
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=> exp	hydroxyethy	lstarch/cn
E1	1	HYDROXYETHYLPACHYMAN/CN
E2	1	HYDROXYETHYLPUERARIN/CN
E3	0>	HYDROXYETHYLSTARCH/CN
E4	1	HYDROXYETHYLTHEOBROMINE/CN
E5	1	HYDROXYETHYLTHEOPHYLLINE/CN
E6	1	HYDROXYETHYLTHIAMINE/CN
E7	1	HYDROXYETHYLTHIAMINEMONOPHOSPHATE/CN
E8	1	HYDROXYETHYLTHIAZOLE KINASE/CN
E9	1	HYDROXYETHYLTHIAZOLE KINASE (4-METHYL-5-BETA-HYDROXYETHYLTHI
		AZOLE KINASE) (LACTOBACILLUS SAKEI SAKEI STRAIN 23K GENE THI

E10	1	M)/CN HYDROXYETHYLTHIAZOLE KINASE (4-METHYL-5-BETA-HYDROXYETHYLTHI AZOLE KINASE) (THZ KINASE) (TH KINASE) (METHANOGENIC ARCHAEO
E11	1	N RICE CLUSTER IMRE50 GENE THIM)/CN HYDROXYETHYLTHIAZOLE KINASE (ACINETOBACTER BAUMANNII STRAIN ATCC 17978)/CN
E12	1	HYDROXYETHYLTHIAZOLE KINASE (ACINETOBACTER STRAIN ADP1 GENE THIM)/CN
=> exp hydro	xvethv	l starch/cn
E1		HYDROXYETHYL PROPYL CELLULOSE/CN
E2	1	HYDROXYETHYL RADICAL/CN
E3	1>	HYDROXYETHYL STARCH/CN
E4	1	HYDROXYETHYL STARCH 130/0.4/CN
E5	1	HYDROXYETHYL STARCH ARGININE ESTER/CN
E6	1	HYDROXYETHYL STARCH LYSINE ESTER/CN
E7	1	HYDROXYETHYL STARCH-POLYETHYLENE GLYCOL DIISOCYANATE COPOLYM ER/CN
E8	1	HYDROXYETHYL STEARYL ETHER/CN
E9	1	HYDROXYETHYL TETRADECYL CELLULOSE/CN
E10	1	HYDROXYETHYL TETRAHYDROPYRANYL CELLULOSE/CN
E11	1	HYDROXYETHYL THIAZOLE KINASE (STAPHYLOCOCCUS AUREUS STRAIN M
		U50 GENE THIM)/CN
E12	1	HYDROXYETHYL THIAZOLE KINASE (STAPHYLOCOCCUS EPIDERMIDIS STR AIN ATCC12228 GENE SE1691)/CN
=> s e3		
L1	1 "HY	DROXYETHYL STARCH"/CN
=> exp hydro	xvet hv	l amylopectin/cn
E1		HYDROXYETHYL ALLYL ETHER-TRIFLUOROCHLOROETHENE-UNDECENOIC AC
		ID-VINYL ACETATE COPOLYMER/CN
E2	1	HYDROXYETHYL AMIDES/CN
E3		HYDROXYETHYL AMYLOPECTIN/CN
E4	1	HYDROXYETHYL AMYLOSE/CN
E5	1	HYDROXYETHYL CARBAMATE/CN
E6	1	HYDROXYETHYL CARBAMATE DIMETHYLOL DIMETHYL ETHER/CN
E7	1	HYDROXYETHYL CARBOXYMETHYL CELLULOSE/CN
E8	1	HYDROXYETHYL CELLULOSE/CN
E9	1	HYDROXYETHYL CELLULOSE 1-OXO-N-OCTADECYL ETHER, 3-SULFO-2-HY
		DROXYPROPYL ETHER/CN
E10	1	HYDROXYETHYL CELLULOSE 2,4-DICHLORO-S-TRIAZIN-6-YL ETHER/CN
E11	1	HYDROXYETHYL CELLULOSE 2-HYDROXY-N-OCTADECYL ETHER, 3-SULFO-
		2-HYDROXYPROPYL ETHER/CN
E12		
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-> 0.62	1	
=> s e3	_	/CN
=> s e3 L2	_	
L2	_	/CN
L2 => s e4	1 "HY	/CN DROXYETHYL AMYLOPECTIN"/CN
L2	1 "HY	/CN
L2 => s e4 L3	1 "HY:	/CN DROXYETHYL AMYLOPECTIN"/CN
L2 => s e4 L3 => file hcap	1 "HY: 1 "HY:	/CN  DROXYETHYL AMYLOPECTIN"/CN  DROXYETHYL AMYLOSE"/CN  SINCE FILE TOTAL
L2 => s e4 L3	1 "HY: 1 "HY:	/CN  DROXYETHYL AMYLOPECTIN"/CN  DROXYETHYL AMYLOSE"/CN  SINCE FILE TOTAL
L2 => s e4 L3 => file hcap	1 "HY 1 "HY lus DOLLA	/CN  DROXYETHYL AMYLOPECTIN"/CN  DROXYETHYL AMYLOSE"/CN  RS  SINCE FILE TOTAL ENTRY SESSION

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FILE COVERS 1907 - 7 Oct 2010 VOL 153 ISS 15
FILE LAST UPDATED: 6 Oct 2010 (20101006/ED)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Aug 2010
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Aug 2010
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HCAplus now includes complete International Patent Classification (IPC) reclassification data for the third quarter of 2010.

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This file contains CAS Registry Numbers for easy and accurate substance identification.

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=> s 11-13
          2824 L1
            12 L2
            24 L3
          2856 (L1 OR L2 OR L3)
T.4
=> s (sterile or sterilization or physiological or intravenous)
         33544 STERILE
         57652 STERILIZATION
         66450 PHYSIOLOGICAL
         45753 INTRAVENOUS
L5
        199427 (STERILE OR STERILIZATION OR PHYSIOLOGICAL OR INTRAVENOUS)
=> s 14 and 15
1.6
           170 L4 AND L5
=> s (molecular weight or mw or dalton or kilodalton or kda or da)
       1475622 MOLECULAR
        196557 WEIGHT
         84326 MOLECULAR WEIGHT
                  (MOLECULAR (W) WEIGHT)
         93687 MW
         11328 DALTON
         16279 KILODALTON
        163908 KDA
         58569 DA
L7
        406196 (MOLECULAR WEIGHT OR MW OR DALTON OR KILODALTON OR KDA OR DA)
=> s 16 and 17
L8
            10 L6 AND L7
=> d 18 1-10 ti abs bib
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- L8 ANSWER 1 OF 10 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Chemically modified konjac glucomannan with high colloid osmotic pressure: Physiological evaluation in a rabbit model as a plasma substitute
- Carboxylmethylated konjac glucomannan (CKGM) is a carboxylmethylated AΒ polymer of mannose and glucose that is derived from the plant Amorphophallus konjac cultivated in East Asia. The CKGM solution had a high volume-expanding efficacy and was evaluated as a plasma substitute in the present study. Ameliorative hemorrhagic shock rabbits were used as the model animals. The in vivo hemodynamic and hemorheol. properties, including blood pressure, blood viscosity, hematocrit, erythrocyte deformation index and erythrocyte aggregation index, were measured in animals treated in the CKGM solution The in vitro colloid osmotic pressure (COP) of the CKGM solution was measured to estimate its plasma-expanding efficacy. These parameters of the CKGM-treated group were compared with groups exposed to 4 other treatments: human serum albumin (HSA), hydroxyethyl starch (HES), polygeline and normal saline. The CKGM solution showed an exceptionally higher COP than other therapy solns. For example, the COP of 1% (weight in volume [w/v]) CKGM solution is comparable to those of

(w/v) HES solution and 5% (w/v) HSA solution Accordingly, the CKGM solution can be

transfused in a much lower dosage while maintaining its plasma-expanding efficacy. The CKGM-treated group showed an improved intravascular persistence and good hemodynamic and hemorheol. properties. Biopsy anal. suggested no organ dysfunction in the group treated in CKGM solution Moreover, the high plasma-expanding efficacy and inexpensive availability of the CKGM solution may facilitate its clin. application as a potential plasma substitute.

- AN 2010:833960 HCAPLUS <<LOGINID::20101007>>
- DN 153:343067

6%

- TI Chemically modified konjac glucomannan with high colloid osmotic pressure: Physiological evaluation in a rabbit model as a plasma substitute
- AU Li, Suping; Hu, Tao; Chen, Yali; Wang, Xianwei; Liu, Tao; Ma, Guanghui; Su, Zhiquo
- CS National Key Laboratory of Biochemical Engineering, Institute of Process Engineering, Beijing, 100190, Peop. Rep. China
- SO Glycobiology (2010), 20(8), 950-958 CODEN: GLYCE3; ISSN: 0959-6658
- PB Oxford University Press
- DT Journal
- LA English
- RE.CNT 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L8 ANSWER 2 OF 10 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Effects of hypertonic-hyperoncotic solution on cardiac function and extravascular lung water in children after open-heart surgery
- AB The effects of hypertonic-hyperoncotic solution (HHS) on cardiac function and extravascular lung water in children after open-heart surgery for congenital cardiac disease were evaluated. Fifty children with congenital cardiac disease were randomly assigned to 2 groups. The HHS group received HHS (7.5% sodium chloride with 6% hydroxyethyl starch 200 kDa). The ISS group (the control group) received isotonic saline solution (ISS 0.9% sodium chloride). Cardiac index (CI), extravascular lung water index (ELWI), stroke volume index (SVI), mean arterial pressure (MAP), and systemic vascular resistance index (SVRI) were measured. Immediately after surgery, patients were loaded either with HHS or with ISS (4 mL/kg). Sodium concentration, osmolality, thrombocyte count (TC), fibrinogen, and arterial blood gases were detected before operation, immediately after loading, 15 min, 1, 4, 12, and 24 h after the end of volume loading. Hemodynamic parameters were recorded at the same time. The total amount of

dobutamine required was documented. In HHS group, MAP, SVI and CI increased, and SVRI decreased significantly after the administration of HHS, compared with ISS group and before administration (P<0.01 or 0.05). Both CVP and HR were unchanged in both groups. In HHS group, ELWI decreased significantly, compared with before volume administration. But ELWI increases directly and remained elevated for 60 min after the administration of ISS. Sodium concentration increased immediately after infusion

of HHS. The postoperative need for infused dobutamine in the patients in HHS group was decreased, compared with ISS group (P<0.05). All patients left the hospital in a clin. sufficient state. A single infusion of HHS after cardiac surgery was safe. After cardiopulmonary bypass surgery, the administration of HHS increased CI by elevating SVI in combination with a decreased SVRI. ELWI significantly decreased, which suggested that HHS effectively counteracted the capillary leakage.

- AN 2009:1145934 HCAPLUS <<LOGINID::20101007>>
- DN 152:165930
- TI Effects of hypertonic-hyperoncotic solution on cardiac function and extravascular lung water in children after open-heart surgery
- AU Li, Danfeng; Wan, Xi; Cheng, Bangchang; Xu, Jinjin
- CS Department of Anaesthesiology, Renmin Hospital, Wuhan University, Wuhan, Hubei Province, 430060, Peop. Rep. China
- SO Zhongguo Yishi Zazhi (2008), 10(12), 1625-1628 CODEN: ZYZHAD; ISSN: 1008-1372
- PB Zhongguo Yishi Zazhishe
- DT Journal
- LA Chinese
- L8 ANSWER 3 OF 10 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Effect of medium-molecular-weight hydroxyethyl starch on the functions of blood coagulation and fibrinolysis
- The study was performed in 30 adult patients, ASA grade I-II, scheduled AB for selective surgery, which were randomly allocated to receiving i.v. infusion of 6% HAES-sterile (HES) group, lactated Ringer's solution (RL), at the rate of  $20mL \cdot kg - 1 \cdot h - 1$  for 60min, resp., with 15 case in each group. Venous blood samples were taken before and after 1h following the infusion to deterion: Hb (Hb), hematocrit (HCT), platelet count (PLT), platelet aggregation test (PAG) [involve PAG(1), PAG(5), PAG(M), T(M)], activated partial thromboplastin time (APTT), prothrombin time (PT), thrombin time (TT), fibrinogen (FIB), tissue plasminogen activator (t-PA), plasminogen activator inhibitor (PAI-1). In HES group, Hb, HCT, PLT, PAG(1) decreased significantly, PT, APTT was prolonged remarkably after infusion, and Hb, HCT, HLT, APTT had marked difference than those in the control group. Following i.v. hydroxyethyl starch, there wre no influences on the circulation and fibrinolysis. HAESsterile 6% had significant effect on intrinsic factors of blood coagulation, but this change was inside the normal range.
- AN 2006:279120 HCAPLUS <<LOGINID::20101007>>
- DN 145:20796
- TI Effect of medium-molecular-weight hydroxyethyl starch on the functions of blood coagulation and fibrinolysis
- AU Xu, Xue; Zhao, Yanli; Jin, Hailong; Cheng, Huiping; Yang, Binxia; Wang, Baoguo
- CS Affiliated Beijing Tiantan Hospital, Capital University of Medical Sciences, Beijing, 100050, Peop. Rep. China
- SO Zhongguo Quanke Yixue (2005), 8(7), 553-555 CODEN: ZQYHAK; ISSN: 1007-9572
- PB Zhongguo Quanke Yixue Zazhishe
- DT Journal
- LA Chinese

- L8 ANSWER 4 OF 10 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Novel pathways in the etiology of cancer, and treatment methods
- AB The invention pertains to the identification of two novel epithelial signaling pathways in ER-pos. breast cancers and the discovery that the cellular biol. and (likely also the clin. outcome) of ER-pos. breast cancer cells is unexpectedly altered when these signaling pathways are activated. The first pathway pertains to the discovery that NF- $\kappa$ B activation and/or DNA binding is implicated in the etiol. of ER-pos. breast (and other) cancers. The second pathway involves ligand-independent quinine-mediated ER activation by phosphorylation (e.g. on SER-118 and SER-167 residues of ER) and nuclear translocation of full-length (67 kDA) ER as well as the phorphorylating activation of a truncated and nuclear-localized ER variant (.apprx.52 kDa). Also disclosed are methods for identifying patients likely to respond to hormonal therapy and for selecting a therapeutic regimen for the treatment of cancer.
- AN 2006:101964 HCAPLUS <<LOGINID::20101007>>
- DN 144:184652
- TI Novel pathways in the etiology of cancer, and treatment methods
- IN Benz, Christopher C.
- PA Buck Institute for Age Research, USA
- SO U.S. Pat. Appl. Publ., 49 pp. CODEN: USXXCO
- DT Patent
- LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI PRAI	US 20060024691 US 2004-556774P US 2004-580534P US 2004-629691P	A1 P P P	20060202 20040325 20040616 20041119	US 2005-90546	20050324

ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT

- L8 ANSWER 5 OF 10 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Pharmaceutical combinations gadolinium complexes and colloidal biopolymers and their use as i.v. injection and i.v. infusion contrast agents in MR angiography
- AB The invention concerns pharmaceutical combinations that contain a low molweight gadolinium complex and a biopolymer in pharmaceutical acceptable carrier; the Gd-complex is in molecularly dispersed form and the biopolymer is colloidally disolved. Gd-DTPA or gadobutrol are formulated with hydroxyethyl starch or acetyl starch. I.v. injections and i.v. infusions for whole body MRI are formulated, especially angiog. and MRI perfusion studies. Field of application is the imaging of cerebrospinal injuries. Thus 10 mL sterile injection solution contained (g): gadopentetic acid dimeglumine salt 4.690; diethylene triamine pentaacetic acid 0.004; meglumine 0.010; hyroxyethyl starach 0.7000; water 6.985.
- AN 2005:822089 HCAPLUS <<LOGINID::20101007>>
- DN 143:199930
- TI Pharmaceutical combinations gadolinium complexes and colloidal biopolymers and their use as i.v. injection and i.v. infusion contrast agents in MR angiography
- IN Tack, Johannes
- PA Neurobiotec G.m.b.H., Germany
- SO Ger. Offen., 7 pp.
  - CODEN: GWXXBX
- DT Patent
- LA German
- FAN.CNT 1

PATENT NO. KIND DATE APPLICATION NO. DATE

DE 102004006048 A1 20050818
DE 102004006048 B4 20091126 \_\_\_\_\_ PΤ DE 2004-102004006048 20040202 PRAI DE 2004-102004006048 20040202 ANSWER 6 OF 10 HCAPLUS COPYRIGHT 2010 ACS on STN L8 ΤI Medicinal agent with volemic effect and method for its preparing The medicinal agent represents hydroxyethylated starch in an aqueous solution AB containing 5-10% of hydroxyethylated starch with the optimal ratio of substituted hydroxyethyl groups at atoms C2/C6 up to 6:1 in glucose residue, average value of mol. mass 130-450 kDa, narrowed mol.-mass distribution at the substitution degree 0.35-0.70 and 0.80-1.00% of sodium chloride. The agent is prepared using maize or potato starch as the raw material with the content of amylopectin 95%, not less. Starch is subjected for alkaline purification, acidic or enzymic hydrolysis up to preparing products with mol. mass 400-900 kDa up to the required degree of substitution of hydroxyethyl groups. The solution is purified from impurities by ultrafiltration and/or reverse osmosis and purification is carried out using apyrogenic activated carbon and/or by sterilizing filtration and the following thermal sterilization of the end product. The invention provides a new agent for rapid blood pressure recovery after blood loss. 2005:120436 HCAPLUS <<LOGINID::20101007>> ΑN DN 142:162697 Medicinal agent with volemic effect and method for its preparing ΤI Panov, V. P.; Korotaev, G. K.; Kir'yanov, N. A.; Panov, A. V.; Dolotov, S. ΙN M.; Leshnevskii, K. A.; Grineva, L. P.; Kotova, Yu. A. PΑ Russia Russ., No pp. given SO CODEN: RUXXE7 DT Patent LA Russian FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE PI RU 2245714 C1 20050210 RU 2003-126930 20030904 PRAI RU 2003-126930 20030904 L8 ANSWER 7 OF 10 HCAPLUS COPYRIGHT 2010 ACS on STN ΤI Blood plasma substitutes containing hydroxyethyl starch with good pH stability, and plastic bags filled with them AΒ Blood plasma substitutes contain hydroxyethyl starch (HES) (Mw 150,000-300,000), Na+ and Cl- as the only electrolytes, and citrate ion as a pH-adjusting agent. A solution (pH 6.5) containing HES (30 g), NaCl (4.5 g), an aqueous 1% Na citrate solution (1.53 mL), and H2O to 100 mL showed pH 5.92 after 30-day storage at 40° after sterilization at 115° for 15 min in a polypropylene bag. 2004:429918 HCAPLUS <<LOGINID::20101007>> ΑN 140:412290 DN TIBlood plasma substitutes containing hydroxyethyl starch with good pH stability, and plastic bags filled with them Tono, Hiroshi; Fujino, Keiichi; Toyama, Toshihiro ΙN PΑ Nihon Pharmaceutical Co., Ltd., Japan Jpn. Kokai Tokkyo Koho, 11 pp. CODEN: JKXXAF DT Patent LA Japanese FAN.CNT 1 KIND DATE APPLICATION NO. DATE PATENT NO. \_\_\_\_\_

PI JP 2004149450 A 20040527 JP 2002-315846 20021030 PRAI JP 2002-315846 20021030

- L8 ANSWER 8 OF 10 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Vaccine composition comprising an immunologically active substance embedded in microparticles consisting of starch with reduced molecular weight
- AB A vaccine composition is disclosed which comprises an immunol. active substance embedded in microparticles essentially consisting of starch having an amylopectin content exceeding 85 % by weight, of which at least 80 % by weight has an average mol. weight within the range of 10-10,000 kDa. A process for preparing such vaccine composition is also disclosed.
- AN 2002:275771 HCAPLUS <<LOGINID::20101007>>
- DN 136:299676
- TI Vaccine composition comprising an immunologically active substance embedded in microparticles consisting of starch with reduced molecular weight
- IN Joensson, Monica; Larsson, Karin; Gustafsson, Nils Ove; Laakso, Timo; Reslow, Mats
- PA Bioglan AB, Swed.
- SO PCT Int. Appl., 61 pp. CODEN: PIXXD2
- DT Patent
- LA English

FAN.CNT 2

FAN.	PAT	CENT I				KIND DATE						DATE								
ΡI										WO 2001-SE2169 200										
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			LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	PH,	PL,		
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	CA	2424	936			A1										20011005				
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		6706			B2	20040316														
	EΡ	1322	290			A1		2003	0702	EP 2001-972895										
		R:				•		ES,		•			LI,	LU,	NL,	SE,	MC,	PT,		
			ΙE,	SI,	LT,	LV,	FΙ,	RO,	MK,	CY,	ΑL,	TR								
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ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS) RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L8 ANSWER 9 OF 10 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Ifosfamide lyophilizate preparations
- The invention relates to improved ifosfamide prepns. which are AB distinguished in that as primary auxiliary a polysaccharide, in general a glycan, preferably dextran, starches or cellulose, in particular dextrans having an MW of 20,000 to 85,000, modified starches such as hydroxyethyl starch and chemical modified celluloses such as hydroxyethylcellulose and sodium CM-cellulose, a glycol ether, preferably polyethylene glycol, in particular polyethylene glycols having a mol. weight of 600 to 6000 or an amino acid, preferably alanine, leucine or glutamic acid, is added to them. The improved ifosfamide preparation can also contain as an auxiliary a pharmaceutically customary buffer, for example acetate, citrate or tris buffer, preferably phosphate buffer. In addition, improved ifosfamide prepns. are obtained by addition of NaHCO3. The ifosfamide prepns. according to the invention can comprise one or a combination of several auxiliaries. Mesna can be added to the formulation as a uroprotector. Ifosfamide 1000 g and alanine 337.3 g were dissolved in 8 L water and the solution was made up to a final volume of 10 L and sterile-filtered. The solution was dispensed under aseptic conditions into sterilized glass vials at 10.0 mL per vial. were transferred to a freeze-drying unit and cooled to a temperature of  $-40^{\circ}$ .

AN 1998:300520 HCAPLUS <<LOGINID::20101007>>

DN 129:8602

OREF 129:1853a,1856a

TI Ifosfamide lyophilizate preparations

IN Wichert, Burkhard; Sauerbier, Dieter; Rawert, Jurgen

PA Asta Medica Aktiengesellschaft, Germany

SO U.S., 4 pp. CODEN: USXXAM

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	US 5750131	A	19980512	US 1996-752069	19961119
PRAI	US 1996-752069		19961119		
ASST	GNMENT HISTORY FOR U	S PATEN'	T AVATLABLE	IN LSUS DISPLAY FORMAT	

OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L8 ANSWER 10 OF 10 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Effects of intravenous pentafraction on lung and soft tissue liquid exchange in hypoproteinemic sheep
- Effects of infusing pentafraction (Pen), a synthetic hydroxyethyl starch AΒ plasma volume expander, on lung and soft tissue lymph flux were compared in nonanesthetized sheep that were protein depleted by batch plasmapheresis. Pen (5%) was infused to raise pulmonary arterial wedge pressure by 5 mmHg for 2 h (1.8). Pen raised plasma osmotic pressure from plasmapheresis baseline (10.7 mmHg; pre-plasmapheresis baseline, 19.6 mmHg) to 16.6 mmHq. After Pen, lung lymph flows peaked at 3.9 times a pre-plasmapheresis baseline value of 1.0 (plasmapheresis baseline, 2.7), but soft tissue lymph flows rose insignificantly. Plasma Pen concns. were 2.3% postinfusion and 1.6% at 12 h. Pen mean mol. masses at these times, measured by high-performance liquid chromatog., were 160 and 129 kDa , resp. In lung lymph, Pen concns. were 0.8% postinfusion and 0.7% at 12h, with mean mol. masses of 125 and 112 kDa, resp. In soft tissue, lymph Pen was nearly undetectable postinfusion, but at 12 h, concns. averaged 0.3% with a mean mol. mass of 80 kDa. The

osmotic effectiveness of Pen may be related to its mol. mass, which was large enough to restrict filtration so that the plasma-to-lung lymph osmotic pressure gradient widened. Pen remained effective in the circulation for at least 24 h. 1994:69160 HCAPLUS <<LOGINID::20101007>> ΑN 120:69160 DN OREF 120:12263a,12266a Effects of intravenous pentafraction on lung and soft tissue liquid exchange in hypoproteinemic sheep Conhaim, R. L.; Rosenfeld, D. J.; Schreiber, M. A.; Baaske, D. M.; Harms, ΑU В. А. CS Dep. Surg., Univ. Wisconsin, Madison, WI, 53705, USA SO American Journal of Physiology (1993), 265(5, Pt. 2), H1536-J1543 CODEN: AJPHAP; ISSN: 0002-9513 DT Journal English LA OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS) => d his log hold 'LOG' IS NOT VALID HERE For an explanation, enter "HELP DISPLAY HISTORY". => d his (FILE 'HOME' ENTERED AT 11:27:18 ON 07 OCT 2010) FILE 'REGISTRY' ENTERED AT 11:27:33 ON 07 OCT 2010 EXP HYDROXYETHYL/CN EXP HYDROXYETHYLSTARCH/CN EXP HYDROXYETHYL STARCH/CN 1 S E3 L1EXP HYDROXYETHYL AMYLOPECTIN/CN 1 S E3 L2L3 1 S E4 FILE 'HCAPLUS' ENTERED AT 11:28:39 ON 07 OCT 2010 L42856 S L1-L3 L5199427 S (STERILE OR STERILIZATION OR PHYSIOLOGICAL OR INTRAVENOUS) L6 170 S L4 AND L5 L7 406196 S (MOLECULAR WEIGHT OR MW OR DALTON OR KILODALTON OR KDA OR DA) L8 10 S L6 AND L7 => log hold COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 54.52 36.82 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE -8.50-8.50

SESSION WILL BE HELD FOR 120 MINUTES
STN INTERNATIONAL SESSION SUSPENDED AT 11:29:59 ON 07 OCT 2010

Connecting via Winsock to STN

## LOGINID: SSPTAEX01623

PASSWORD:

\* \* \* \* \* RECONNECTED TO STN INTERNATIONAL \* \* \* \* \* SESSION RESUMED IN FILE 'HCAPLUS' AT 12:02:08 ON 07 OCT 2010 FILE 'HCAPLUS' ENTERED AT 12:02:08 ON 07 OCT 2010 COPYRIGHT (C) 2010 AMERICAN CHEMICAL SOCIETY (ACS)

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 36.82 54.52

SINCE FILE DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) TOTAL SESSION ENTRY

CA SUBSCRIBER PRICE -8.50-8.50

=> s 11/thu or 12/thu or 13/thu

2824 L1

1300368 THU/RL

1083 L1/THU

(L1 (L) THU/RL)

12 L2

1300368 THU/RL

1 L2/THU

(L2 (L) THU/RL)

24 L3

1300368 THU/RL

1 L3/THU

(L3 (L) THU/RL)

L9 1085 L1/THU OR L2/THU OR L3/THU

=> s 17 and 19

102 L7 AND L9 L10

=> file stnguide

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 39.73 57.43

DISCOUNT AMOUNTS (FOR OUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION

CA SUBSCRIBER PRICE -8.50-8.50

FILE 'STNGUIDE' ENTERED AT 12:02:47 ON 07 OCT 2010 USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT COPYRIGHT (C) 2010 AMERICAN CHEMICAL SOCIETY (ACS)

FILE CONTAINS CURRENT INFORMATION.

LAST RELOADED: Oct 1, 2010 (20101001/UP).

=> fiel hcaplus

FIEL IS NOT A RECOGNIZED COMMAND

The previous command name entered was not recognized by the system. For a list of commands available to you in the current file, enter "HELP COMMANDS" at an arrow prompt (=>).

=> file hcaplus

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST 0.07 57.50

SINCE FILE TOTAL
ENTRY SESSION
0.00 -8.50

CA SUBSCRIBER PRICE

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FILE COVERS 1907 - 7 Oct 2010 VOL 153 ISS 15 FILE LAST UPDATED: 6 Oct 2010 (20101006/ED) REVISED CLASS FIELDS (/NCL) LAST RELOADED: Aug 2010 USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Aug 2010

HCAplus now includes complete International Patent Classification (IPC) reclassification data for the third quarter of 2010.

CAS Information Use Policies apply and are available at:

http://www.cas.org/legal/infopolicy.html

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 110 and (PY<2006 or AY<2006 or PRY<2006) 26341581 PY<2006 5556094 AY<2006 5045948 PRY<2006

L11 69 L10 AND (PY<2006 OR AY<2006 OR PRY<2006)

=> s intravenous or plasma or (volume expander) or (degree of substitution)

45753 INTRAVENOUS

1084989 PLASMA

142479 VOLUME

3937 EXPANDER

31 VOLUME EXPANDER

(VOLUME (W) EXPANDER)

4810767 DEGREE

310750 SUBSTITUTION

7971 DEGREE OF SUBSTITUTION

(DEGREE (1W) SUBSTITUTION)

L12 1130548 INTRAVENOUS OR PLASMA OR (VOLUME EXPANDER) OR (DEGREE OF SUBSTIT UTION)

=> s 111 and 112

L13 42 L11 AND L12

=> d 113 1-42 ti abs bib

- L13 ANSWER 1 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI An improved process for producing pyrogen-free narrow molecular weight distribution tetrastarch as a plasma volume expander
- AB An improved process for for producing pyrogen-free tetrastarch, used as a plasma volume expander, involves hydrolyzing the waxy starch to achieve a mol. weight distribution of 110,000-150,000 daltons. A molar substitution of 0.35-0.45 is achieved by attachment of hydroxyethyl group. An organic solvent used till tetrastarch becomes free from salt and glycol content. Microfiltration is used to reduce microorganism load (bio-burden) and ultrafiltration to reduce pyrogen and low as well as high mol. weight undesired fragments. A spray dryer is used to obtain dried tetrastarch with the min. moisture content.
- AN 2009:808651 HCAPLUS <<LOGINID::20101007>>
- DN 152:554662
- TI An improved process for producing pyrogen-free narrow molecular weight distribution tetrastarch as a plasma volume expander
- PA Claris Lifesciences Limited, India
- SO Indian Pat. Appl., 15pp.

CODEN: INXXBQ

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	IN 2004MU00423	A	20090626	IN 2004-MU423	20040408 <
PRAI	IN 2004-MU423		20040408	<	

- L13 ANSWER 2 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Effects of PentaLyte and Voluven hemodilution on plasma coagulation kinetics in the rabbit: role of thrombin-fibrinogen and factor XIII-fibrin polymer interactions
- AΒ Background: Hydroxyethyl starch (HES) administration has resulted in decreased hemostasis and fibrinogen (F1)-thrombin-(FIIa)-Factor XIII (FXIII) interactions. I proposed to determine the hemostatic effect of hemodilution with PentaLyte (6% HES, mean mol. weight 220 kDa) and Voluven (6% HES, 130 kDa). Methods: Rabbits were i.v. administered 20 mL/kg PentaLyte or Voluven (n = 8 per fluid) over 10 min. Plasma was obtained prior to, 1 min and 1 h after hemodilution. Thrombelastog. was performed, with clot initiation (R, sec), clot propagation  $(\alpha, \text{ degrees})$ , and clot strength (shear elastic modulus, G, dynes/cm2) determined over 20 min. Celite-activated samples had either no addns. or addition of FI, FIIa or activated FXIII (FXIIIa) to restore protein content to pre-diluted values. Results and conclusions: While there were no significant differences between the groups, R significantly decreased 1 h after hemodilution compared with values observed before and 1 min after hemodilution, whereas  $\alpha$  and G significantly decreased 1 min after hemodilution and then significantly, but only partially, increased 1 h after hemodilution compared with pre-dilution values. Addition of FI, FIIa and FXIIIa significantly decreased R in both groups.  $\alpha$  And G 1 min after hemodilution were significantly enhanced by FI, FIIa, FXIIIa in both groups; however, 1 h after hemodilution, rabbits administered PentaLyte had  $\alpha$  and G enhanced only by FI and FXIIIa addition, whereas animals administered Voluven had  $\alpha$  and G significantly enhanced by FI addition PentaLyte and Voluven hemodilution initially diminishes FIIa-FI and FXIIIa-fibrin, but within an hour primarily inhibit FXIIIa-fibrin interactions in the rabbit.
- AN 2005:1158830 HCAPLUS <<LOGINID::20101007>>
- DN 144:163874
- TI Effects of PentaLyte and Voluven hemodilution on plasma

```
coagulation kinetics in the rabbit: role of thrombin-fibrinogen and factor
     XIII-fibrin polymer interactions
ΑU
    Nielsen, V. G.
     Department of Anesthesiology, The University of Alabama at Birmingham,
CS
     Birmingham, AL, USA
     Acta Anaesthesiologica Scandinavica (2005), 49(9), 1263-1271
SO
     CODEN: AANEAB; ISSN: 0001-5172
PΒ
     Blackwell Publishing Ltd.
DT
     Journal
     English
LA
OSC.G
       12
              THERE ARE 12 CAPLUS RECORDS THAT CITE THIS RECORD (12 CITINGS)
              THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 25
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L13 ANSWER 3 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
     Production and use of hydroxyethyl starch
TI
     Hydroxyethyl starch (I) useful in pharmaceuticals and having weight-average
AΒ
mol.
     weight (Mn) \geq500,000, degree of substitution (DS)
     0.25-0.5, and C2/C6 ratio 2-<8 is prepared Suspending 30 kg wax-cornstarch
     in 52.2 kg H2Oactivating at 85° with 5.1 g NaOH, adding 4.159 kf
     liquid ethylene oxide, heating slowly to 40^{\circ}, stirring for 2 h,
     reducing Mn by heating with 20% HCl (giving pH 2.0) at 75°, cooling
     to 50°, and ultrafiltration gave I with DS 0.39, Mw 1520,
     and C2-C6 ratio 2.3. Use of I as, i.a., a plasma volume expander
     is exemplified.
     2005:979667 HCAPLUS <<LOGINID::20101007>>
ΑN
DN
    143:250014
ΤI
    Production and use of hydroxyethyl starch
    Boll, Michael; Fisch, Andreas; Spahn, Donat R.
ΤN
     B. Braun Melsungen A.-G., Germany
PA
     PCT Int. Appl., 39 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     German
FAN.CNT 1
                   KIND DATE
                                                                 DATE
    PATENT NO.
                                          APPLICATION NO.
                     A2 20050909 WO 2005-EP50877
A3 20060316
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    WO 2005082942
                                                                  20050301 <--
     WO 2005082942
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
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             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
             NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM,
             SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
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             RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
             MR, NE, SN, TD, TG
                                20050909
                                            AU 2005-217157
     AU 2005217157
                         Α1
                                                                   20050301 <--
                                           EP 2005-708068
     EP 1732953
                         Α2
                                20061220
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     EP 1732953
                               20071107
                         В1
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             IS, IT, LI, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR
     CN 1926155
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                               20070307
                                            CN 2005-80006765
                                                                   20050301 <--
    BR 2005008285 A 20070807
JP 2007525588 T 20070906
AT 377609 T 20071115
ZA 2006008126 A 20080227
                                            BR 2005-8285
                                                                   20050301 <--
                                          JP 2007-501278
                                                                  20050301 <--
                                         AT 2005-708068
                                                                  20050301 <--
                                           ZA 2006-8126
                                                                   20050301 <--
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ES 2294680
                      T3 20080401 ES 2005-708068
C2 20091120 RU 2006-134639
                     Т3
                                                              20050301 <--
    RU 2373222
                                                              20050301 <--
    IN 2006CN03159
                      A
                             20070608 IN 2006-CN3159
                                                             20060831 <--
                                       KR 2006-7020430
    KR 2007022672
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                             20070227
                                                              20060929 <--
    US 20070282014
                      A1
                            20071206
                                       US 2007-590462
                                                              20070730 <--
PRAI EP 2004-100813
                      Α
                            20040301 <--
    WO 2005-EP50877
                       W
                             20050301 <--
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ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT RE.CNT 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L13 ANSWER 4 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Albumin and hydroxyethyl starch 130 kDa/0.4 improve filter clearance and hemocompatibility in hemo- and plasmafiltration--an in vitro study
- Apart from their standard applications, hemofiltration (HF) and plasma AΒ filtration (PF) may provide helpful therapy for sepsis, multiple organand acute liver-failure. Some colloids cause either decreases or increases in blood cell agglomeration. We hypothesized that solns. which reduce cell aggregability may lead to both improved filter clearance and better haemocompatibility due to decreasing rates of clogged hollow fibers. Heparinized porcine blood (5 IU/mL) was used in an in vitro circuit. The filter types tested were from GABBRO: HF66D (effective membrane surface: 0.6 m2) and PF1000N (effective membrane surface: 0.15 m2). Albumin (ALB), hydroxyethyl starch (HES) 200/0.5, HES 130/0.4, gelatin (GEL) or normal saline (0.9%) were added to the blood (n = 6/group). Recirculation systems were run for 2 h. Spontaneous hemollysis and filter resistance >420 mmHg were selected as indications of maximal flow rates. Sieving coeffs. were determined for 17 parameters at the lowest and highest blood flows and filtration rate. Based on the filter types used, supplementation of ALB and HES130/0.4 led to an improved filter clearance without increasing the number of clogged capillary membranes or causing impaired haemocompatibility. Sieving coeffs. for most solutes were independent of volume substitute and flow rate. Haemocompatibility and filter clearance deteriorated after addition of  ${\tt HES200}$  or  ${\tt GEL}$  to the blood. Under standardized in vitro conditions, we found that colloids which reduce cell aggregability cause improved HF- and PF-performance. This phenomenon may provide new options for higher clearances and may lead to new concepts in low dose anticoagulation.
- AN 2005:886981 HCAPLUS <<LOGINID::20101007>>
- DN 144:494927
- TI Albumin and hydroxyethyl starch 130 kDa/0.4 improve filter clearance and hemocompatibility in hemo- and plasmafiltration--an in vitro study
- AU Unger, Juliane K.; Haltern, Claudia; Dohmen, Bernd; Gressner, Axel; Grosse-Siestrup, Christian; Groneberg, David A.; Rossaint, Rolf
- CS Department of Anaesthesiology, University Hospital Aachen, Rheinisch-Westfalische Technische Hochschule Aachen, Aachen, Germany
- SO Nephrology, Dialysis, Transplantation (2005), 20(9), 1922-1931 CODEN: NDTREA; ISSN: 0931-0509
- PB Oxford University Press
- DT Journal
- LA English
- OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)
  RE.CNT 20 THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD
  ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 5 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Antithrombin efficiency is maintained in vitro in human plasma following dilution with hydroxyethyl starches
- AB Hemodilution has been associated with changes in hemostasis secondary to

modulation of procoagulant activity. However, direct effects of specific fluids on anticoagulants, such as antithrombin (AT), remained undefined. Thus, the purpose of this investigation was to determine whether hemodilution with hydroxyethyl starches (HES) directly diminishes plasma AT activity, which would be manifested by decreases in clot initiation time (reaction time, R) with thrombelastog. greater than that seen with 0.9% NaCl (NS). Normal plasma and AT-deficient (< 1% activity) plasma were diluted 0 or 30% with NS, Hextend (6% HES; average mol. weight, 450 kDa), PentaLyte (6% HES; average mol. weight, 220 kDa), or Voluven (6% HES; average mol. weight, 130 kDa) (n = 6-7 expts. per condition). Undiluted, normal plasma had an R value of  $796 \pm$ 65 s, which was significantly (P < 0.05) greater than R values following NS (690  $\pm$  50 s) or Voluven (675  $\pm$  68 s) dilution R values of normal plasma diluted with Hextend (831  $\pm$  51 s) or PentaLyte (801  $\pm$ 72 s) were not different from undiluted plasma but were significantly (P < 0.05) greater than those observed following NS or Voluven dilution There were no significant differences between the conditions when AT-deficient plasma was utilized (R range, 404-440 s). Rather than interfere with AT activity, HES with an average mol. weight of 220-450 kDa maintain AT efficiency.

- AN 2005:535082 HCAPLUS <<LOGINID::20101007>>
- DN 143:359705
- TI Antithrombin efficiency is maintained in vitro in human plasma following dilution with hydroxyethyl starches
- AU Nielsen, Vance G.
- CS Department of Anesthesiology, The University of Alabama at Birmingham, Birmingham, AL, 35249-6810, USA
- SO Blood Coagulation & Fibrinolysis (2005), 16(5), 319-322 CODEN: BLFIE7; ISSN: 0957-5235
- PB Lippincott Williams & Wilkins
- DT Journal
- LA English
- OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)
- RE.CNT 18 THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 6 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Molecular weight of hydroxyethyl starch: is there an effect on blood coagulation and pharmacokinetics?
- AΒ Background: The development of hydroxyethyl starches (HES) with low impact on blood coagulation but higher volume effect compared with the currently used HES solns. is of clin. interest. We hypothesized that high mol. weight, low-substituted HES might possess these properties. Methods: Thirty pigs were infused with three different HES solns. (20 mL kg-1) with the same degree of molar substitution (0.42) but different mol. wts. (130, 500) and 900 kDa). Serial blood samples were taken over 24 h and blood coagulation was assessed by Thromboelastograph anal. and anal. of plasma coagulation. In addition, plasma concentration and in vivo mol. weight were determined and pharmacokinetic data were computed based on a two-compartment model. Results: Thromboelastograph anal. and plasma coaquiation tests did not reveal a more pronounced alteration of blood coagulation with HES 500 and HES 900 compared with HES 130. In contrast, HES 500 and HES 900 had a greater area under the plasma concentration-time curve [1542 (142) g min litre-1, P<0.001, 1701 (321) g min litre-1, P<0.001] than HES 130 [1156 (223) g min litre-1] and alpha half life ( $t\alpha 1/2$ ) was longer for HES 500 [53.8 (8.6) min, P<0.01] and HES 900 [57.1 (12.3) min, P<0.01] than for HES 130 [39.9 (10.7) min]. Beta half life (t $\beta$ 1/2), however, was similar for all three types of HES [from 332 (100) to 381 (63) min]. Conclusions: In low-substituted HES, mol. weight is not a key factor in compromising blood coagulation. The longer initial intravascular persistence of high mol.

- weight low-substituted HES might result in a longer lasting volume effect.
- 2005:312645 HCAPLUS <<LOGINID::20101007>> ΑN
- DN 143:146205
- Molecular weight of hydroxyethyl starch: is there an ΤI effect on blood coagulation and pharmacokinetics?
- Madjdpour, C.; Dettori, N.; Frascarolo, P.; Burki, M.; Boll, M.; Fisch, ΑU A.; Bombeli, T.; Spahn, D. R.
- Department of Anaesthesiology, University Hospital Lausanne, Lausanne, CS CH-1011, Switz.
- SO British Journal of Anaesthesia (2005), 94(5), 569-576 CODEN: BJANAD; ISSN: 0007-0912
- РΒ Oxford University Press
- DTJournal
- LA English
- THERE ARE 16 CAPLUS RECORDS THAT CITE THIS RECORD (16 CITINGS) OSC.G 16
- RE.CNT 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- ANSWER 7 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN L13
- Medicinal agent with volemic effect and method for its preparing ТΤ
- AΒ The medicinal agent represents hydroxyethylated starch in an aqueous solution containing 5-10% of hydroxyethylated starch with the optimal ratio of substituted hydroxyethyl groups at atoms C2/C6 up to 6:1 in glucose residue, average value of mol. mass 130-450 kDa, narrowed mol.-mass distribution at the substitution degree 0.35-0.70 and 0.80-1.00% of sodium chloride. The agent is prepared using maize or potato starch as the raw material with the content of amylopectin 95%, not less. Starch is subjected for alkaline purification, acidic or enzymic hydrolysis up to

preparing

products with mol. mass 400-900 kDa up to the required degree of substitution of hydroxyethyl groups. The solution is purified from impurities by ultrafiltration and/or reverse osmosis and purification is carried out using apyrogenic activated carbon and/or by sterilizing filtration and the following thermal sterilization of the end product. The invention provides a new agent for rapid blood pressure recovery after blood loss.

- 2005:120436 HCAPLUS <<LOGINID::20101007>> ΑN
- DN 142:162697
- ΤI Medicinal agent with volemic effect and method for its preparing
- Panov, V. P.; Korotaev, G. K.; Kir'yanov, N. A.; Panov, A. V.; Dolotov, S. M.; Leshnevskii, K. A.; Grineva, L. P.; Kotova, Yu. A.
- PΑ Russia
- SO Russ., No pp. given CODEN: RUXXE7
- Pat.ent. DΤ
- LA Russian
- FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ΡI	RU 2245714	C1	20050210	RU 2003-126930	20030904 <
PRAI	RU 2003-126930		20030904	<	

- ANSWER 8 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- Infusion solutions containing polymers for the diagnosis and therapy of ΤI
- The invention concerns infusion solns. containing non-lipophilic, biol. inert macromols. that (a) have mol. wts. of 6-60 kDa; (b) become enriched in tumors because of the altered permeability of tumor vessels;
  - (c) are selected from the group of plasma volume expanders; and
  - (d) there can be small mols. coupled to the macromols. Applied macromols. are gelatin, polysuccinate, hydroxyethyl starch, dextran, inulin,

oxypolygelatin, crosslinked polypeptides, polyhydroxyethyl aspartamide and their mixture. The concentration of the macromols. is typically 6-10%.

agents and anticancer agents can be coupled to the polymers.

AN 2004:956508 HCAPLUS <<LOGINID::20101007>>

DN 141:415976

- TI Infusion solutions containing polymers for the diagnosis and therapy of tumors
- PA Tritthart, Helmut A., Austria

SO Austrian, 5 pp.

CODEN: AUXXAK

DT Patent

LA German

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE					
ΡI	AT 409929	В	20021227	AT 1997-387	19970307 <					
	AT 9700387	A	20020515							
PRAI	AT 1997-387		19970307	<						

- L13 ANSWER 9 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI The Effects of High Molecular Weight Hydroxyethyl Starch Solutions on Platelets
- AB Physicochem. characteristics of hydroxyethyl starch (HES) mols. determine their side effects on hemostasis. Our aim in the present expts. was to test the antiplatelet effect of novel high mol. weight HES. Citrated whole blood was hemodiluted in vitro (0% and 20%) with either HES 550 (Hextend), HES 600 (6%Hetastarch-Baxter), HES 200 (Elohaest), or the solvent of Hextend in its com. available solution The availability of glycoprotein IIb-IIIa was assessed on nonstimulated and on agonist-induced platelets using flow cytometry. Glycoprotein IIb-IIIa availability increased significantly after hemodilution with Hextend and its solvent by 23% and 24%, resp., but decreased in the presence of 6% Hetastarch-Baxter and Elohaest by 18% and 15%, resp., with no significant difference between the latter two colloids. This study shows that Hextend does not inhibit platelet function as anticipated by its high mol. weight and degree of substitution. The unexpected platelet stimulating effect of Hextend is unique among the currently available HES prepns. and may, at least in part, be induced by its solvent containing calcium chloride dihydrate (2.5 mmol/L). The platelet-inhibiting effect of 6%Hetastarch-Baxter was not significantly different from that of medium mol. weight HES 200.
- AN 2004:679330 HCAPLUS <<LOGINID::20101007>>
- DN 142:245773
- TI The Effects of High Molecular Weight Hydroxyethyl Starch Solutions on Platelets
- AU Deusch, Engelbert; Thaler, Ulrich; Kozek-Langenecker, Sibylle A.
- CS Department of Anesthesiology and Intensive Care, Vienna Medical University, Austria
- SO Anesthesia & Analgesia (Hagerstown, MD, United States) (2004), 99(3), 665-668
- CODEN: AACRAT; ISSN: 0003-2999
  PB Lippincott Williams & Wilkins
- DT Journal
- LA English
- OSC.G 18 THERE ARE 18 CAPLUS RECORDS THAT CITE THIS RECORD (18 CITINGS)
- RE.CNT 25 THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 10 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Blood plasma substitutes containing hydroxyethyl starch with good pH stability, and plastic bags filled with them

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Blood plasma substitutes contain hydroxyethyl starch (HES) (
AΒ
     Mw 150,000-300,000), Na+ and Cl- as the only electrolytes, and
     citrate ion as a pH-adjusting agent. A solution (pH 6.5) containing HES (30
g),
     NaCl (4.5 g), an aqueous 1% Na citrate solution (1.53 mL), and H2O to 100 mL
     showed pH 5.92 after 30-day storage at 40° after sterilization at
     115° for 15 min in a polypropylene bag.
     2004:429918 HCAPLUS <<LOGINID::20101007>>
ΑN
DN
     140:412290
     Blood plasma substitutes containing hydroxyethyl starch with
ΤI
     good pH stability, and plastic bags filled with them
     Tono, Hiroshi; Fujino, Keiichi; Toyama, Toshihiro
ΙN
PA
     Nihon Pharmaceutical Co., Ltd., Japan
SO
     Jpn. Kokai Tokkyo Koho, 11 pp.
     CODEN: JKXXAF
DT
     Patent
     Japanese
LA
FAN.CNT 1
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     PATENT NO.
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     JP 2004149450
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                                   20040527
PRAI JP 2002-315846
                                   20021030 <--
    ANSWER 11 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
     Highly-branched, low substituted starch products for use as plasma
TI
     expanders
     The invention concerns modified hydroxyethyl and hydroxypropyl starches
AB
     for clin. use as plasma expanders that have a branching degree
     of 8-20 mol%, a substitution degree (MS) of 0.05-0.3 and mol. weight of
     10,000-450,000. The products are used in peritoneal dialysis. According
     to expts. with rats, the products deplete faster from liver, spleen, lung
     and kidney than conventional starch products.
     2004:198158 HCAPLUS <<LOGINID::20101007>>
ΑN
DN
     140:223241
ΤI
     Highly-branched, low substituted starch products for use as plasma
     expanders
ΙN
     Henning, Klaus
     Fresenius Kabi Deutschland G.m.b.H., Germany
PA
     Ger. Offen., 5 pp.
     CODEN: GWXXBX
DT
     Patent
LA
     German
FAN.CNT 1
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     DE 10237442
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PΙ
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                           В4
     DE 10237442
                                   20040819
     WO 2004022602 A1 20040318
                                                                    20030730 <--
                                              WO 2003-EP8411
          W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
              CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
         PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
              BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                       A1 20040329 AU 2003-251668 20030730 <--
A1 20050518 EP 2003-793660 20030730 <--
     AU 2003251668
     EP 1530593
          R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
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IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
    CN 1675248
                              20050928
                                         CN 2003-819356
                       Α
                                                               20030730 <--
    CN 100340578
                        С
                              20071003
    JP 2005539107
                        Т
                              20051222
                                         JP 2004-533291
                                                               20030730 <--
    US 20060032400
                           20060216 US 2005-524424
                                                               20050722 <--
                       A1
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                       B2 20090623
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                       A1 20080627
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                       А
                                                               20100521 <--
PRAI DE 2002-10237442
                       Α
                            20020816 <--
                           20030730 <--
    JP 2004-533291
                        АЗ
    WO 2003-EP8411
                        W
                             20030730 <--
ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT
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             THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)
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THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 2 ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L13 ANSWER 12 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- ΤI Volume efficacy and reduced influence on measures of coagulation using hydroxyethyl starch 130/0.4 (6%) with an optimised in vivo molecular weight in orthopaedic surgery: a randomised, double-blind study
- AΒ Background and objective: Different types of hydroxyethyl starch (HES) affect blood coaqulation differently. The authors studied the effects of HES 130/0.4 on coagulation in major orthopedic surgery in relation to the pharmacol. parameter in vivo mol. weight Methods: 52 patients were randomly allocated to either HES 130/0.4 (6%, mean mol. weight 130 kDa, molar substitution 0.4) or HES 200/0.5 (6%, control) in a double-blind fashion. Colloidal volume requirements for intra- and postoperative hemodynamic stabilization were compared. Safety analyses of this pharmacol. study included a comparison of coagulation factor tests, in vivo mol. weight, and HES plasma concns. Results: The colloidal vols. given were similar at the end of surgery (1602±569 for HES  $130/0.4 \text{ vs. } 1635\pm567 \text{ mL for HES } 200/0.5), 5 \text{ h later } (1958 \pm,467 \text{ vs.})$  $1962\pm398$  mL), and up to the first postoperative day ( $2035\pm446$  vs.  $2000\pm424$  mL). HES in vivo mol. weight at the end of surgery was  $88,707\pm13$  938 vs.  $158,374\pm33$  933 Da (p < 0.001) and 5 h later was  $86,663\pm16$  126 vs.  $136,299\pm26$  208 Da (p < 0.001). In parallel to the lower in vivo mol. weight, factor VIII and von Willebrand factor returned to almost normal in the HES 130/0.4 group up to 5 h postoperatively, but not in the control group (p < 0.05) Residual HES plasma concns. after 24 h were low in the HES 130/0.4 group (1.0 mq/mL), but higher in the control group (2.6 mg/mL). Conclusion: HES 130/0.4 and HES 200/0.5 were found to be similar with regard to volume efficacy. Sensitive coagulation parameters returned more rapidly to normal in the HES 130/0.4 group. Lower in vivo mol. weight and more rapid excretion of HES 130/0.4 are the likely explanations for the smaller influence on coagulation in this group.
- 2004:179543 HCAPLUS <<LOGINID::20101007>> ΑN
- DN 140:228979
- Volume efficacy and reduced influence on measures of coagulation using TIhydroxyethyl starch 130/0.4 (6%) with an optimised in vivo molecular weight in orthopaedic surgery: a randomised, double-blind study
- Jungheinrich, Cornelius; Sauermann, Wilhelm; Bepperling, Frank; Vogt, ΑU Norbert H.
- CS Clinical Research, Fresenius Kabi, Bad Homburg, Germany
- SO Drugs in R&D (2004), 5(1), 1-9CODEN: DRDDFD; ISSN: 1174-5886
- ΡВ Adis International Ltd.
- DT Journal
- LA English

- OSC.G 23 THERE ARE 23 CAPLUS RECORDS THAT CITE THIS RECORD (23 CITINGS)
  RE.CNT 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD
  ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 13 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Binding of hydroxyethyl starch molecules to the platelet surface
- AΒ Hydroxyethyl starch (HES) solns. impair platelet function by reducing the availability of the fibrinogen receptor. This effect is not mediated by intracellular signal transduction pathways. Also, an unspecific coating of platelets by HES macromols. may be responsible for its antiplatelet effects. To test this hypothesis, the authors investigated the binding of fluorochrome-coupled HES to the surface of human platelets using whole blood flow cytometry. Citrated whole blood from 8 volunteers was incubated (5 min, 22°C, in the dark) with fluorescein isothiocyanate (FITC)-coupled HES (200-kDa mol. weight, 0.5 degree of substitution, 0.042 molar ratio of FITC-conjugation) resulting in 0, 1, 3, 5, 10, 20, and 40% hemodilution. The percentage of platelets binding FITC-HES was determined using a FACSCalibur flow cytometer and CellQuestPro software. The percentage of FITC-pos. platelets increased in a concentration-dependent manner reaching statistical significance at 10% hemodilution. Binding was independent of fibrinogen receptor blockade. The present expts. clearly demonstrate that extracellular binding of HES to the platelet surface is, at least in part, responsible for the antiplatelet effects of HES by blocking the access of ligands to the platelet fibrinogen receptor.
- AN 2003:759942 HCAPLUS <<LOGINID::20101007>>
- DN 140:139068
- TI Binding of hydroxyethyl starch molecules to the platelet surface
- AU Deusch, Engelbert; Gamsjager, Thomas; Kress, Hans-Georg; Kozek-Langenecker, Sibylle A.
- CS Department of Anesthesiology and Intensive Care (B), School of Medicine, University of Vienna, Vienna, Austria
- SO Anesthesia & Analgesia (Hagerstown, MD, United States) (2003), 97(3), 680-683 CODEN: AACRAT; ISSN: 0003-2999
- PB Lippincott Williams & Wilkins
- DT Journal
- LA English
- OSC.G 21 THERE ARE 21 CAPLUS RECORDS THAT CITE THIS RECORD (21 CITINGS)
- RE.CNT 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 14 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Influence of long-term volume therapy with hydroxyethyl starch on leukocytes in patients with acute stroke
- A repeated administration of hydroxyethyl starch affects hemostasiol. and AΒ rheol. factors such as the concentration of factor VIII/von Willebrand factor, platelet volume and plasma viscosity. An earlier study showed that HES also lowers the concentration of fibronectin, a mol. important in the reticulo-endothelial system (RES). RES has a "clearing function" and is a part of the non-immune-specific defense mechanisms of the body. It is involved in the elimination of HES from the blood. Since leukocytes are another important part of the unspecific defense system, the goal of the present study was to investigate whether HES affects leukocytes. After giving their informed consent, 20 patients with cerebral perfusion disorders were randomized and underwent a double-blind 10-day hypervolemic hemodilution with HES 200/0.5/13 or HES 70/0.5/4. The nos. of leukocytes, percentage of lymphocytes, percentage of neutrophilic granulocytes and Hb concentration were measured. The absolute number of leukocytes did not change significantly, but the share of neutrophilic granulocytes increased. The increase in neutrophilic granulocytes reflects an increase in phagocytic

activity. HES 200/0.5/13, which has the larger in vivo mol. weight ( MW = 95 kD), caused a larger increase in neutrophilic granulocytes than HES 70/0.5/4, which has an in vivo MW of 58 kD.

- AN 2003:549860 HCAPLUS <<LOGINID::20101007>>
- DN 139:224104
- TI Influence of long-term volume therapy with hydroxyethyl starch on leukocytes in patients with acute stroke
- AU Woessner, Ralph; Grauer, Markus T.; Dieterich, Hans-Juergen; Treib, Wolfgang; Stoll, Martin; Treib, Johannes
- CS Neurologische Klinik, Westpfalz-Klinikum GmbH, Kaiserslautern, Germany
- SO Arzneimittel-Forschung (2003), 53(6), 402-406 CODEN: ARZNAD; ISSN: 0004-4172
- PB Editio Cantor Verlag
- DT Journal
- LA English
- OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)
- RE.CNT 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 15 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI The effects of hydroxyethyl starch solutions on thromboelastography in preoperative male patients
- AΒ Hydroxyethyl starches (HES) have been shown to decrease clot strength and to increase coaquiation times assessed by thromboelastoq. (TEG). HES with minimal anticoagulant side-effects is beneficial for plasma volume expansion in the perioperative setting. A comparison of the in vivo effects of high, middle and low mol. weight HES solns. on TEG variables has not been performed so far. Blood was obtained before and after i.v. infusion (10 mL kg-1) of either saline, HES 70/0.5/4 (mol. weight in kDa/degree of substitution/C2:C6 ratio), HES 130/0.4/9, HES 200/0.6/9.4, or HES 450/0.7/4.6 in 50 otherwise healthy patients. Thromboelastog. was performed in 360  $\mu l$  of 1% celite activated citrated whole blood after recalcification. HES 450/0.7/4.6 prolonged reaction time indicating impairment of the plasmatic coagulation system. TEG parameters indicative for platelet function, including angle  $\alpha$ , maximum amplitude and coagulation time, deteriorated after infusion of HES 450/0.7/4.6 and HES 70/0.5/4. HES 200/0.6/9.4 and HES 130/0.4/9impaired platelet contribution to hemostasis only partially, decreasing two or one TEG platelet parameters, resp. Infusion of HES 450/0.7/4.6 compromises TEG parameters more than the other solns. tested, whereas HES 130/0.4/9 has the smallest effect. Further outcome-related studies are needed to assess the clin. relevance of our findings.
- AN 2003:137114 HCAPLUS <<LOGINID::20101007>>
- DN 138:297342
- TI The effects of hydroxyethyl starch solutions on thromboelastography in preoperative male patients
- AU Felfernig, M.; Franz, A.; Braunlich, P.; Fohringer, C.; Kozek-Langenecker, S. A.
- CS Department of Anesthesiology and Intensive Care B, School of Medicine, University of Vienna, Vienna, Austria
- SO Acta Anaesthesiologica Scandinavica (2003), 47(1), 70-73 CODEN: AANEAB; ISSN: 0001-5172
- PB Blackwell Munksgaard
- DT Journal
- LA English
- OSC.G 13 THERE ARE 13 CAPLUS RECORDS THAT CITE THIS RECORD (13 CITINGS)
- RE.CNT 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 16 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Hydroxyethyl starch does not cross the blood-brain or the placental

- barrier but the perineurium of peripheral nerves in infused animals Therapy with hydroxyethyl starch (HES) is associated with a high incidence of AΒ persistent pruritus due to HES storage in cutaneous nerves. Up to now it has been unknown if HES also accumulates in the extra-cutaneous peripheral or central nervous system. To study this, five rats including one pregnant one were infused with a single dose (34-150 mg) of HES (70/200/450 kDa mol. weight) conjugated with fluorescein isothiocyanate (FITC). In addition, four sheep were infused with a cumulative dosage of 30 g, 120 g, and 420 g HES (200 kDa), resp. After 7-13 days, biopsies from the adult rats, four fetal rats and sheep were taken from various organs. The specimens were analyzed by light, electron, and confocal laser scanning microscopy. Typical HES storage vacuoles were found in macrophages of the skin, liver, spleen, lung, and kidney. HES storage in healthy animals was not associated with signs of either inflammation or apoptosis contrary to a previously described animal hemorrhagic shock model. Beyond that, fetus biopsies did not show any storage phenomenon, confirming that HES does not cross the placental barrier. Deposits of HES could be detected in Schwann cells of cutaneous nerve fibers as well as in perineural and endoneural cells of sciatic nerve in one rat (HES 450 kDa) and three of four sheep. No HES storage was found in the central nervous system. Our findings clearly demonstrate that storage of HES is detectable only in small peripheral nerves, suggesting a cutaneous origin of the HES-induced pruritus.
- AN 2003:86491 HCAPLUS <<LOGINID::20101007>>
- DN 139:143288
- TI Hydroxyethyl starch does not cross the blood-brain or the placental barrier but the perineurium of peripheral nerves in infused animals
- AU Stander, S.; Bone, H. G.; Machens, H. G.; Aberle, T.; Burchard, W.; Prien, T.; Luger, T. A.; Metze, D.
- CS Department of Dermatology, University of Munster, Munster, 48149, Germany
- SO Cell & Tissue Research (2002), 310(3), 279-287 CODEN: CTSRCS; ISSN: 0302-766X
- PB Springer-Verlag
- DT Journal
- LA English
- OSC.G 5 THERE ARE 5 CAPLUS RECORDS THAT CITE THIS RECORD (5 CITINGS) RE.CNT 40 THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS RECORD
  - ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 17 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Hydroxyethylstarch as a risk factor for acute renal failure: is a change of clinical practice indicated?
- A review. Hypovolemia is extremely common among surgical and intensive AB care patients. The best strategy for volume replacement therapy has been the focus of debate for several years. The lack of acceptance of hydroxyethylstarch (HES) for volume replacement therapy is most likely due to reports of abnormal coagulation and to recently published studies indicating neg. effects of HES on renal function. All HES solns. are not created equal - they widely differ with regard to their physicochem. characteristics (concentration, mean mol. weight (Mw), degree of substitution [DS], C2/C6-substitution ratio). These differences have important consequences for adverse effects such as alterations in the coagulation process and on kidney function. Conflicting results about the effects of different HES solns. on renal function may also be due to varying clin. protocols, selection of patients, and different criteria for volume replacement. Theor. and documented hazards are associated with each kind of volume replacement therapy. There appears to be no reason to banish modern HES prepns. with a low or medium Mw (e.g. 70, 130 or 200kD) and a low DS (0.4 or 0.5) in patients without pre-existing kidney dysfunction. In patients with known renal dysfunction (e.g. plasma creatinine level >3 mg/dL), all HES prepns. should be used

cautiously and other volume replacement regimens (e.g. gelatins) should be considered since no convincing data are yet available for the latest generation of HES (Mw 130; DS 0.4).

- AN 2002:856392 HCAPLUS <<LOGINID::20101007>>
- DN 137:345468
- TI Hydroxyethylstarch as a risk factor for acute renal failure: is a change of clinical practice indicated?
- AU Boldt, Joachim
- CS Department of Anaesthesiology and Intensive Care Medicine, Klinikum der Stadt Ludwigshafen, Ludwigshafen, Germany
- SO Drug Safety (2002), 25(12), 837-846 CODEN: DRSAEA; ISSN: 0114-5916
- PB Adis International Ltd.
- DT Journal; General Review
- LA English
- OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)
- RE.CNT 42 THERE ARE 42 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 18 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Effect of plasma for volume replacement in induced hypovolemic shock. A comparative study of low and medium molecular weight hydroxyethyl starch substitutes
- AB We studied the relative efficacy of two plasma substitute therapies in a canine model for hemorrhagic shock. Anesthetized dogs were bled to maintain a mean arterial pressure (mAP) at 50 mmHg and then administered a single bolus injection of 6% hydroxyethyl starch (HES) with a mol. weight of 70 kDa (HES70 group) or 200 kDa (HES200 group) at a volume equivalent to the blood withdrawn. The efficacy of both therapies in maintaining the hemodynamic variables, the plasma colloidal and crystalloidal osmotic pressure (Pcop and Posm), and the circulating blood volume (CBV) were investigated. CBV was measured by the pulse-dye densitometry (PDD) method. After resuscitation, hemodynamic variables were better maintained in the HES200 group than in the HES70 group. Particularly, mAP, mean pulmonary arterial pressure, pulmonary arterial wedge pressure, cardiac index, left ventricular stroke work index, and maximum rate of left ventricular pressure change, were maintained at a satisfactorily stable level in the HES200 group as compared with the HES70 group. Moreover, Pcop and CBV in the HES200 group were significantly greater than those in the HES70 group. On the other hand, Posm did not differ between the two groups. HES200 may be a more effective volume replacement therapy than HES70 for induced hemorrhagic shock because of improvement and maintenance of hemodynamic variables, CBV and Pcop.
- AN 2002:791203 HCAPLUS <<LOGINID::20101007>>
- DN 137:315846
- TI Effect of plasma for volume replacement in induced hypovolemic shock. A comparative study of low and medium molecular weight hydroxyethyl starch substitutes
- AU Maruta, Kyoko
- CS Dep. Anesthesiol., Sch. Med. Showa Univ., Japan
- SO Showa Igakkai Zasshi (2002), 62(3), 188-193 CODEN: SIGZAL; ISSN: 0037-4342
- PB Showa Daigaku, Showa Igakkai
- DT Journal
- LA Japanese
- L13 ANSWER 19 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Pharmacodynamics and tolerability of acetyl starch as a new plasma volume expander in patients undergoing elective surgery
- AB Acetyl starch (ACS) is a new synthetic colloid solution for plasma

volume expansion and is now undergoing phase II clin. trials. We compared the pharmacodynamics and tolerability of ACS with those of hydroxyethyl starch (HES) in 32 patients (American Society of Anesthesiologists phys. status I and II) undergoing elective surgery. In this prospective, randomized, double-blind trial patients received either 15 mL/kg ACS 6% (average mol. weight (Mw) 200,000/ molar substitution (MS) 0.5) or HES 6% (Mw 200,000/ MS 0.5) i.v. up to a maximum dose of 1000 mL. Hemodynamic parameters, rheol. parameters, volume effect, acid-base status as well as effects on hemostasis were studied. After infusion of ACS and HES there was a similar increase in central venous pressure and mean arterial pressure in both groups. Acid-base status was not significantly altered after the end of the colloid infusions. After ACS infusion, plasma acetate concentration increased from 0.13±0.16 mg/dL to 2.87±1.13 mg/dL, however, after 24 h there was no significant difference in plasma acetate concentration compared to HES. The volume effect ranged from 104-116% (ACS) and from 88-118% (HES) of the colloid dose administered. These differences were not statistically significant. Partial thromboplastin time (aPTT) was only slightly increased after ACS infusion (from  $38.6\pm5.7$  s to  $41.4\pm5.1$  s), but was significantly increased after HES infusion (from 38.7±5.7 s to 46.1±7.0 s). ACS and HES are equally effective plasma volume expanders; ACS might be a new, alternative colloid solution with fewer coagulation side-effects than HES.

- AN 2002:88807 HCAPLUS <<LOGINID::20101007>>
- DN 136:272953
- TI Pharmacodynamics and tolerability of acetyl starch as a new plasma volume expander in patients undergoing elective surgery
- AU Bremerich, D. H.; Lischke, V.; Asskali, F.; Forster, H.; Behne, M.
- CS Department of Anesthesiology and Resuscitation,
  Johann-Wolfgang-Goethe-Universitatsklinikum, Frankfurt, Germany
- SO International Journal of Clinical Pharmacology and Therapeutics (2000), 38(8), 408-414 CODEN: ICTHEK; ISSN: 0946-1965
  - Dustri-Verlag Dr. Karl Feistle
- DT Journal

PΒ

- LA English
- OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)
  RE.CNT 18 THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD
  ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 20 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Effects of resuscitation with hydroxyethyl starch (HES) on pulmonary hemodynamics and lung lymph balance in hemorrhagic sheep; comparative study of low- and high-molecular-weight HES
- AB Studies of extremely low- and high-mol.-weight HES were performed to evaluate the effects of these solns. on lung lymph filtration during resuscitation. Conscious sheep were bled from an arterial line to maintain shock. After 2 h of hemorrhage, the following solns. were infused for 1 h: low-mol.-weight HES (mol. weight 70,000, substitution fractions 0.5-0.55); high-mol.-weight HES (mol. weight 450,000, substitution fractions 0.65); normal saline. The amount of solution infused was the same as the volume of blood lost. Both low- and high-mol.-weight HES equally restored systemic arterial pressure and cardiac output and increased pulmonary microvascular pressure. However, the actual oncotic pressure gradient (plasma/lymph) rose transiently during infusion of low-mol.-weight HES, while high-mol.-weight HES increased the

oncotic pressure gradient even after cessation of the infusion. Lung lymph flow during and after resuscitation with low-mol.-weight HES and saline rose significantly from the preshock value. There was no significant difference between low-mol.-weight HES and saline with respect to effects on lung lymph flow. However, lung lymph flow after high-mol.-weight HES was

less than that after low-mol.-weight HES. These data suggest that low-mol.-weight HES is as useful as a plasma substitute as high-mol.-weight HES but has the possibility of increasing lung lymph filtration during the early phase of resuscitation.

- AN 2002:41206 HCAPLUS <<LOGINID::20101007>>
- DN 137:195249
- TI Effects of resuscitation with hydroxyethyl starch (HES) on pulmonary hemodynamics and lung lymph balance in hemorrhagic sheep; comparative study of low- and high-molecular-weight HES
- AU Kaneki, Toshimichi; Koizumi, Tomonobu; Yamamoto, Hiroshi; Fujimoto, Keisaku; Kubo, Keishi; Shibamoto, Toshishiqe
- CS First Department of Internal Medicine, Shinshu University School of Medicine, Shinshu, 390-8621, Japan
- SO Resuscitation (2002), 52(1), 101-108 CODEN: RSUSBS; ISSN: 0300-9572
- PB Elsevier Science Ireland Ltd.
- DT Journal
- LA English
- OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)
- RE.CNT 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 21 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Effects of colloidal resuscitation fluids on the neutrophil respiratory burst
- Exptl. studies have revealed that gelatin and HES produce increased AΒ neutrophil respiratory burst activity. It was investigated whether 3-percent gelatin (MW 35,000) and three types of 6-percent HES ( MW 70,000; degree of substitution, 0.5; 200,000/0.5; 450,000/0.7) prepns. can influence superoxide anion production during respiratory burst under clin. conditions. Blood samples were obtained from 40 patients before and 1 h after the infusion, before anesthesia and surgical treatment. After stimulation with bacteria (Escherichia coli), the respiratory burst was measured by oxidation of non-fluorescent dihydrorhodamine 123 to the fluorescent rhodamine 123 by the use of flow cytometry. Respiratory burst activity decreased significantly (p = 0.004) from the baseline (60.0 ± 6.5%) to 1 h after the administration of the low-mol.-weight HES preparation (55.0  $\pm$  6.8%). No significant differences in respiratory burst activity could be found after the administration of gelatin or medium-mol.-weight or high-mol.-weight HES solution Thus, the administration of gelatin and medium- and high-mol.-weight HES prepns. did not influence respiratory burst activity under clin. conditions. However, the neutrophil respiratory burst was impaired after the administration of low-mol.-weight HES. Neutrophil respiratory burst activity may vary according to the type of colloidal plasma substitutes administered.
- AN 2001:648196 HCAPLUS <<LOGINID::20101007>>
- DN 136:334969
- TI Effects of colloidal resuscitation fluids on the neutrophil respiratory burst
- AU Jaeger, Karsten; Heine, Joern; Ruschulte, Heiner; Juttner, Bjorn; Scheinichen, Dirk; Kuse, Ernst R.; Piepenbrock, Siegfried
- CS Departments of Anesthesiology and Intensive Care Medicine and of Abdominal and Transplantation Surgery, Hannover Medical School, Hannover, D-30625, Germany
- SO Transfusion (Bethesda, MD, United States) (2001), 41(8), 1064-1068

  CODEN: TRANAT; ISSN: 0041-1132
- PB American Association of Blood Banks
- DT Journal
- LA English

- OSC.G 9 THERE ARE 9 CAPLUS RECORDS THAT CITE THIS RECORD (9 CITINGS)
  RE.CNT 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD
  ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 22 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Effects of hydroxyethylstarch and gelatin on renal function in severe sepsis: a multicenter randomized study
- Hydroxyethylstarch used for volume restoration in brain-dead kidney donors AΒ has been associated with impaired kidney function in the transplant recipients. We undertook a multicenter randomized study to assess the frequency of acute renal failure (ARF) in patients with severe sepsis or septic shock treated with hydroxyethylstarch or gelatin. Adults with severe sepsis or septic shock were enrolled prospectively in three intensive-care units in France. They were randomly assigned 6% hydroxyethylstarch (200 kDa, 0.60-0.66substitution) or 3% fluid-modified gelatin. The primary endpoint was ARF (a two-fold increase in serum creatinine from baseline or need for renal replacement therapy). Analyses were by intention to treat. Severity of illness and serum creatinine (median 143 [IQR 88-203] vs. 114 [91-175]  $\mu$ mol/L) were similar at baseline in the hydroxyethylstarch and gelatin groups. The frequencies of ARF (27/65 [42%] vs. 15/64 [23%], p=0.028) and oliquria (35/62 [56%] vs. 23/63 [37%], p=0.025) and the peak serum creatinine concentration (225 [130-339] vs. 169 [106-273]  $\mu$ mol/L, p=0·04) were significantly higher in the hydroxyethylstarch group than in the gelatin group. In a multivariate anal., risk factors for acute renal failure included mech. ventilation (odds ratio 4.02 [95% CI 1.37-11.8], p=0.013) and use of hydroxyethylstarch (2.57 [1.13-5.83], p=0.026). The use of this preparation of hydroxyethylstarch as a plasma-volume expander is an independent risk factor for ARF in patients with severe sepsis or septic shock.
- AN 2001:221381 HCAPLUS <<LOGINID::20101007>>
- DN 135:220973
- TI Effects of hydroxyethylstarch and gelatin on renal function in severe sepsis: a multicenter randomized study
- AU Schortgen, F.; Lacherade, J.-C.; Bruneel, F.; Cattaneo, I.; Hemery, F.; Lemaire, F.; Brochard, L.
- CS Medical Intensive-Care Unit, Hopital Henri Mondor, Assistance Publique-Hopitaux de Paris, University Paris 12, Creteil, 94000, Fr.
- SO Lancet (2001), 357(9260), 911-916 CODEN: LANCAO; ISSN: 0140-6736
- PB Lancet Ltd.
- DT Journal
- LA English
- OSC.G 58 THERE ARE 58 CAPLUS RECORDS THAT CITE THIS RECORD (58 CITINGS)
- RE.CNT 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 23 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI The influence of intravascular volume therapy with a new hydroxyethyl starch preparation (6% HES 130/0.4) on coagulation in patients undergoing major abdominal surgery
- AB A new hydroxyethyl starch (HES) preparation with a mean mol. weight of 130,000 Da and a degree of substitution of 0.4 shows favorable pharmacokinetic properties. We conducted a study of the influence of the new HES specification on coagulation and compared it with another colloidal intravascular volume replacement regimen using gelatin. According to a prospective, random sequence, 42 patients undergoing major abdominal surgery received either HES 130/0.4 (n = 21) or gelatin (n = 21) until the first postoperative day (POD) to keep central venous pressure between 10 and 14 mm Hg. From arterial blood samples, standard coagulation

variables were measured, and modified thromboelastogram (TEG) measurements using different activators were performed. A total of 2830±350 mL of gelatin and  $2430\pm310$  mL of HES 130/0.4 were administered until the morning of the first POD. The use of allogeneic blood/blood products and standard coagulation variables did not differ significantly between the two groups. After induction of anesthesia, all TEG data for both groups were within normal range. Coaqulation time and maximum clot firmness did not change significantly in any TEG measurements during the study period. The kinetics of clot formation (clot formation time) significantly increased immediately after surgery, but without showing significant group differences. On the morning of the first POD, the clot formation time returned to almost normal levels, except for aprotinin-activated TEG. We conclude that administration of moderate doses of the new HES 130/0.4preparation in patients undergoing major abdominal surgery results in similar coagulation alterations as those after using an established gelatin-based volume-replacement regimen.

- AN 2001:219315 HCAPLUS <<LOGINID::20101007>>
- DN 135:174970
- TI The influence of intravascular volume therapy with a new hydroxyethyl starch preparation (6% HES 130/0.4) on coagulation in patients undergoing major abdominal surgery
- AU Haisch, Gerd; Boldt, Joachim; Krebs, Claudia; Kumle, Bernhard; Suttner, Stefan; Schulz, Andreas
- CS Department of Anesthesiology and Intensive Care Medicine, Klinikum der Stadt Ludwigshafen, Ludwigshafen, D-67063, Germany
- SO Anesthesia & Analgesia (Baltimore, MD, United States) (2001), 92(3), 565-571 CODEN: AACRAT; ISSN: 0003-2999
- PB Lippincott Williams & Wilkins
- DT Journal
- LA English
- OSC.G 35 THERE ARE 35 CAPLUS RECORDS THAT CITE THIS RECORD (35 CITINGS)
- RE.CNT 26 THERE ARE 26 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 24 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Influence of colloid fluids on polymorphonuclear granulocyte function in vivo
- AΒ Granulocytes have a role in the immediate immune response. In a previous investigation the authors could demonstrate in vitro a moderate increase of the complement receptors CR1 (CD35) and CR3 (CD11b/CD18) on the surface of polymorphonuclear neutrophils (PMN) after incubation of whole blood with colloids. To elucidate the clin. significance, the authors investigated if these changes were also present in vivo. The study was performed prior to anesthesia for orthopedic surgery. A total of 60 ASA-I patients was evaluated. Patients received in a randomized manner 7 mL/kg of the following solns.: human albumin 5% (HA), gelatine 4% (GEL), hydroxyethylstarch solution 6% with MW 200 000 Da, degree of substitution 0.5 (HES), or Ringer's solution Prior to the infusion, at the end (30 min) and again 30 min later, blood samples were taken. Blood was incubated with fluorescein-conjugated monoclonal antibodies (CD11b, CD16, CD35, CD62L) and analyzed with flow cytometry. HA, GEL, HES, and Ringer's solution failed to induce significant differences in the expression of complement receptors CR1 (CD35) and CR3 (CD11b/CD18), Fc $\gamma$  receptor IIIb (CD16), and of L-selectin (CD62L) receptor on the surface of PMN. Application of colloids like HA, GEL, or HES in moderate amts. shows no short-term effect on adhesion or activation mols. on granulocytes. However, in high doses, infused in situations such as multiple trauma and sepsis, the consequences on the function of PMN may be speculative and require further investigations.
- AN 2001:212429 HCAPLUS <<LOGINID::20101007>>

- DN 135:200293
- TI Influence of colloid fluids on polymorphonuclear granulocyte function in vivo
- AU Engel, J. M.; Welters, I.; Rupp, M.; Langefeld, T.; Ruwoldt, R.; Menges, T.; Hempelmann, G.
- CS Department of Anaesthesiology and Intensive Care Medicine, Justus-Liebig-University, Giessen, Germany
- SO Acta Anaesthesiologica Scandinavica (2001), 45(3), 385-389 CODEN: AANEAB; ISSN: 0001-5172
- PB Munksquard International Publishers Ltd.
- DT Journal
- LA English
- OSC.G 11 THERE ARE 11 CAPLUS RECORDS THAT CITE THIS RECORD (11 CITINGS)
- RE.CNT 25 THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 25 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Low- and medium-molecular-weight hydroxyethyl starches: Comparison of their effect on blood coagulation
- AB High-mol.-weight hydroxyethyl starch (HES) compromises blood coagulation more than medium-mol.-weight HES. The authors compared medium mol. weight HES (200 kd [HES200]) and low-mol.-weight HES (70 kd [HES70]). In a prospective, double-blind, randomized-sequence crossover study, 22 male volunteers received 15 mL/kg HES200 and HES70. Blood samples were taken before and 5 min, 30 min, 1 h, 2 h, 4 h, 8 h, and 24 h after infusion. The following parameters were analyzed at all time points: prothrombin time, activated partial thromboplastin time, fibrinogen, factor VIII, antigenetic and functional von Willebrand factor, platelets, Thrombelastograph anal. parameters (reaction time, coagulation time, maximum amplitude, angle  $\alpha$ , and clot lysis 30 and 60 min after maximum amplitude), ionized Ca, hematocrit, HES blood plasma concentration, mol. weight (weight average and number

average), molar substitution, and polydispersity (weight average/number average).

Repeated-measures anal. of variance was used to compare the response of the aforementioned parameters to the infusion of HES70 and HES200. Both HES solns, had an impact on all parameters. A slightly greater compromise with HES200 was found in activated partial thromboplastin time, factor VIII, antigenetic von Willebrand factor, functional von Willebrand factor, maximum amplitude, and angle  $\alpha$ . No difference was established with the other parameters. HES concentration, weight average, number average, and polydispersity were

higher with HES200. There was no difference with molar substitution. Low-mol.-weight hydroxyethyl starch (70 kd) compromises blood coagulation slightly less than HES200, but it is unclear whether this is clin. relevant.

- AN 2000:850186 HCAPLUS <<LOGINID::20101007>>
- DN 135:302
- TI Low- and medium-molecular-weight hydroxyethyl starches: Comparison of their effect on blood coagulation
- AU Jamnicki, Marina; Bombeli, Thomas; Seifert, Burkhardt; Zollinger, Andreas; Camenzind, Vladimir; Pasch, Thomas; Spahn, Donat R.
- CS Institute of Anesthesiology, University Hospital, Zurich, CH-8091, Switz.
- SO Anesthesiology (2000), 93(5), 1231-1237 CODEN: ANESAV; ISSN: 0003-3022
- PB Lippincott Williams & Wilkins
- DT Journal
- LA English
- OSC.G 31 THERE ARE 31 CAPLUS RECORDS THAT CITE THIS RECORD (31 CITINGS)
- RE.CNT 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L13 ANSWER 26 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Plasma substituents for volume replacement in hemorrhagic shock: comparison of low and medium molecular weight hydroxyethyl starch
- The aim of this study was to assess the relative efficacy of 2 volume AB replacement therapies in a canine model of induced hemorrhagic shock. Anesthetized dogs were bled to maintain mean arterial pressure (mAP) at 50 mm Hq for 30 min and then administered a single bolus injection of 6% hydroxyethyl starch (HES) with a mol. weight of 70 kd (HES70 group) or 200 kd (HES200 group) at a volume equivalent to the blood withdrawn. The authors examined the efficacy of both therapies in maintaining hemodynamic variables and splanchnic organ blood flow (ie, blood flow through the renal cortex, renal medulla, liver, and pancreas). After resuscitation, hemodynamic variables were better maintained in the HES200 group than in the HES70 group. In particular, HES200 better preserved mAP, cardiac index, mean pulmonary arterial pressure, pulmonary arterial wedge pressure, left ventricular stroke work index, and maximum rate of left ventricular pressure change. In both groups splanchnic organ blood flows decreased after hemorrhagic shock but increased after volume replacement resuscitation. After resuscitation splanchnic organ blood flow was greater in the HES200 group than in the HES70 group. The results of this study suggest that HES200 is more effective than HES70 as volume replacement therapy in a canine model of hemorrhagic shock, as measured by improvements in hemodynamic variables and splanchnic organ blood flow.
- AN 2000:598288 HCAPLUS <<LOGINID::20101007>>
- DN 134:65969
- TI Plasma substituents for volume replacement in hemorrhagic shock: comparison of low and medium molecular weight hydroxyethyl starch
- AU Kobori, Masao; Negishi, Hideru; Nagai, Hiroe; Iyama, Kyoko
- CS Department of Anesthesiology, Showa University School of Medicine, Tokyo, 142-8666, Japan
- SO Current Therapeutic Research (2000), 61(7), 414-421 CODEN: CTCEA9; ISSN: 0011-393X
- PB Excerpta Medica, Inc.
- DT Journal
- LA English
- RE.CNT 14 THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 27 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI First human studies with a high-molecular-weight iron chelator
- The release of free, reactive iron from cellular iron stores has been AΒ implicated as an important contributor to tissue damage in a variety of clin. situations, including ischemia and reperfusion injury, hemorrhagic shock, and burn injury. Deferoxamine mesylate (DFO), the only iron chelator currently approved for clin. use, is used for the treatment of iron overload, including acute iron poisoning and treatment of chronic iron overload in transfusion-dependent anemias such as  $\beta$ -thalassemia. However, it is not suitable for acute care situations because of its toxicity, primarily hypotension when given at high i.v. doses, and its short plasma half-life. We have produced a high-mol.-weight iron chelator by chemical coupling DFO to hydroxyethyl starch. This novel chelator (HES-DFO) was administered to healthy male subjects by i.v. infusion over a 4-h period. The drug was well tolerated, and signs of DFO acute toxicity were not observed Maximum plasma chelator levels of approx. 3 mmol/L were achieved with HES-DFO, which is more than an order of magnitude higher than has been reported with injections of DFO. Drug residence time in plasma was markedly prolonged, with an initial

half-life of 22 to 33 h. Urinary iron excretion was  $7.1 \pm 2.2$  mg in 48 h in the highest dose group, as compared with  $0.06 \pm 0.15$  mg in control subjects who received normal saline infusions. I.v. infusion of HES-DFO is well tolerated, produces substantial and prolonged plasma chelator levels, and markedly stimulates urinary iron excretion.

- AN 2000:430799 HCAPLUS <<LOGINID::20101007>>
- DN 133:37931
- TI First human studies with a high-molecular-weight iron chelator
- AU Dragsten, Paul R.; Hallaway, Philip E.; Hanson, Gregory J.; Berger, Arthur E.; Bernard, Bruce; Hedlund, Bo E.
- CS Biomedical Frontiers Inc, Minneapolis, MN, 55414, USA
- SO Journal of Laboratory and Clinical Medicine (2000), 135(1), 57-65
  - CODEN: JLCMAK; ISSN: 0022-2143
- PB Mosby, Inc.
- DT Journal
- LA English
- OSC.G 24 THERE ARE 24 CAPLUS RECORDS THAT CITE THIS RECORD (24 CITINGS)
- RE.CNT 55 THERE ARE 55 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 28 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI A study of plasma substitutes for volume replacement in intraoperative hemodilution technique ''estimation of circulating blood volume by pulse dve-densitometry
- The purpose of this study was to exptl. compare the hemodynamic variables, AB plasma colloidal and crystalloid osmotic pressure (Pcop and Posm), and circulating blood volume (CBV), under normovolemic hemodilution in isoflurane-anesthetized dogs. We divided anesthetized dogs into two groups: a HES 70 group (hydroxyethyl starch, MW=70 kDa , 6% in saline), and a HES 200 groups (hydroxyethyl starch, MW =200 kDa, 6% in saline). Hemodilution was produced by exchanging blood (25mL/kg) with isovolemic artificial colloid of either HES 70 or HES 200. Measurements and sampling were taken before hemodilution, at the end of hemodilution, and 30, 60, 120, 180, 240, and 300 min after hemodilution. CBV was measured by pulse-dye densitometry (PDD) method. A significant increase in mean pulmonary arterial pressure (mPAP), cardiac index (CI), left ventricular stroke work index (LVSWI), and maximum rate of left ventricular pressure change (LV dp/dt maximum), and a significant decrease in systemic vascular resistance (SVR) values occurred after hemodilution in both groups. However, mAP, mPAP, pulmonary artery wedge pressure (PAWP) and LV dp/dt maximum values in group HES 70 decreased significantly over time compared with the pre-hemodilution condition. MPAP, CI, LVAWI and LV dp/dt maximum values in group HES 200 increased significantly. After hemodilution, CBV and Pcop increased significantly compared with the pre-hemodilution condition in both groups. In group HES 70, CBV and Pcop decreased from the pre-hemodilution condition over time, but not in group HES 200. Moreover, CBV and Pcop in group HES 200 significantly greater than those in group HES 70. Posm did not change significantly during any of the exptl. periods compared to the pre-hemodilution condition in both groups. These results suggest that HES 200 may be more effective than HES 70 for the normovolemic hemodilution. This is due to an improvement and a maintenance in hemodynamic variables, CBV and Pcop.
- AN 2000:403485 HCAPLUS <<LOGINID::20101007>>
- DN 133:256612
- TI A study of plasma substitutes for volume replacement in intraoperative hemodilution technique ''estimation of circulating blood volume by pulse dye-densitometry
- AU Nagai, Hiroe; Kobori, Masao

- CS School of Medicine, Showa University, The purpose of this study was to experimentally compare the, Japan
- SO Junkan Seigyo (2000), 21(1), 47-53 CODEN: JUSEE7; ISSN: 0389-1844
- PB Nippon Junkan Seigyo Igakkai
- DT Journal
- LA English
- RE.CNT 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 29 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Randomized trial of hydroxyethyl starch versus gelatine for trauma resuscitation
- AB Previous studies have demonstrated the rapid increase in systemic capillary permeability after blunt trauma and its association with poor outcome. There are theor. advantages in resuscitation with colloid fluids, which are well retained in the vascular compartment during times of capillary leak. The aim of this study was to compare the effects of post-trauma resuscitation with hydroxyethyl starch (HES) (mol. mass, 250 kDa) or gelatine (mol. mass, 30 kDa), the hypothesis being that HES would reduce capillary leak. Forty-five patients suffering blunt trauma were randomized on admission to receive either gelatine (Gelofusine) (n = 21) or HES (Pentaspan) (n = 24) for the first 24 h, after which the choice of fluid was at the discretion of the clinician. The mean Injury Severity Score for the HES and gelatine groups were 20.0 (range, 9-41) and 18.1 (range, 9-32), resp. (p = 0.43). Capillary permeability was assessed by urine albumin excretion rate for the first 24 h. For 5 days the daily mean PO2/FIO2 ratio, serum C-reactive protein, Hb, white cell and platelet counts, prothrombin, and activated partial thromboplastin time were recorded. Capillary permeability was lower in HES-treated patients during the first 24 h. Log mean (95% confidence interval) albumin excretion rate for gelatine and HES groups at 6 h were 117.5 (84.9) and 46.8 (24.3)  $\mu$ g/min (p = 0.011), at 12 h were 54.9 (30.0) and 17.2 (7.6)  $\mu$ g/min (p = 0.001), and at 24 h were 50.5 (23.4) and 23.6 (16.3)  $\mu g/min$  (p = 0.030), resp. The mean (95% confidence interval) PO2/FIO2 ratio for the HES and gelatine groups 48 h after admission were 324 (44) and 267 (43) mm Hg, resp. (p = 0.03). The mean (95% confidence interval) serum C-reactive protein in the HES and gelatine groups 24 h after admission were 72.4 (19.2) and 105.7 (30.1) mg/L, resp. (p = 0.03). There were no significant differences in any of the hematol. parameters during the first 48 h. The results suggest that compared with gelatine, resuscitation with HES reduces posttrauma capillary leak.
- AN 2000:37173 HCAPLUS <<LOGINID::20101007>>
- DN 132:102622
- TI Randomized trial of hydroxyethyl starch versus gelatine for trauma resuscitation
- AU Allison, Keith P.; Gosling, Peter; Jones, Sarah; Pallister, Ian; Porter, Keith M.
- CS West Midlands Regional Training Scheme, Solihull, West Midlands, B911TA, UK
- SO Journal of Trauma: Injury, Infection, and Critical Care (1999), 47(6), 1114-1121 CODEN: JOTRFA; ISSN: 1079-6061
- PB Lippincott Williams & Wilkins
- DT Journal
- LA English
- OSC.G 22 THERE ARE 22 CAPLUS RECORDS THAT CITE THIS RECORD (22 CITINGS)
- RE.CNT 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 30 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN

- TI Hydroxyethylstarch conjugates their production, and contrast agents containing them
- AB Injected conjugates of hydroxyethylstarch with metal complexes remain confined to the intravascular space and are therefore useful as blood pool contrast agents in medical diagnosis. These agents accumulate in regions with high vascular permeability such as tumors, and can be used to demonstrate the degree of tissue perfusion, e.g. in diagnosis of myocardial infarction. They show high relaxivity in MRI, and have a carrying capacity for paramagnetic ions of .apprx.20%. They show good excretion behavior, good stability, and good biocompatibility (no data). Thus, hydroxyethylstarch (mol. weight 40 kDa) reacted with C1CH2CO2H in alkaline solution to form Na O-(carboxymethyl)hydroxyethylstarch (degree of substitution 1.1), which was amidated with the Gd complex of 10-(2-hydroxy-3-aminopropyl)-4,7,10-tris(carboxymethyl)-1,4,7,10-tetraazacyclododecane.
- AN 1999:549187 HCAPLUS <<LOGINID::20101007>>
- DN 131:185191
- TI Hydroxyethylstarch conjugates their production, and contrast agents containing them
- IN Mareski, Peter; Platzek, Johannes; Raduechel, Bernd; Niedballa, Ulrich;
  Weinmann, Hanns-Joachim
- PA Schering Aktiengesellschaft, Germany
- SO PCT Int. Appl., 34 pp. CODEN: PIXXD2
- DT Patent
- LA German
- FAN.CNT 1

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- L13 ANSWER 31 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Effects of hydroxyethyl starch infusion on lung fluid balance in hemorrhagic sheep
- AB The present study was designed to investigate the effect of relatively low mol. hydroxyethyl starch (HES:Mw 70,000) on pulmonary hemodynamics and lymph flow balance during resuscitation from hemorrhagic hypotension employing instrumented and unanesthetized sheep with chronic lung lymph fistula. After baseline measurements for 2 h, animals were bled from a catheter placed in the artery to maintain systemic hypotension of 60-65 mmHg. After establishment of hemorrhagic hypotension, HES (HES group: n = 6) or normal saline (NS group: n = 5) was infused for one hour. The volume of infused solution was equal to the volume of shed blood in each animal. HES infusion restored systemic arterial pressure much more rapidly than NS. HES also produced significant increases in pulmonary arterial and left atrial pressures, and cardiac output. These parameters at the end of HES infusion were significantly higher than those with NS.

The actual oncotic pressure gradient (plasma-lymph) was transiently widened during HES infusion. Both HES and NS infusion produced an increase in lung lymph flow, but these increased levels did not show significant differences (4.8±1.6 mL/15 min with HES vs. 3.8±1.2 mL/15 min with NS). In conclusion, low mol. HES is a useful plasma substitute as it produced a transient beneficial effect on the oncotic gradient in pulmonary hemodynamics during the resuscitation from hemorrhage. HES solution also did not cause extravascular water retention that might induce respiratory disturbance at the early stage of resuscitation from hemorrhagic hypovolemia.

- AN 1999:447446 HCAPLUS <<LOGINID::20101007>>
- DN 131:134472
- TI Effects of hydroxyethyl starch infusion on lung fluid balance in hemorrhagic sheep
- AU Kaneki, Toshimichi
- CS Sch. Med., Shinshu Univ., Matsumoto, 390-8621, Japan
- SO Shinshu Igaku Zasshi (1999), 47(2), 119-128 CODEN: SIZAA7; ISSN: 0037-3826
- PB Shinshu Igakkai
- DT Journal
- LA Japanese
- L13 ANSWER 32 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Characterization of acetyl starch by means of NMR spectroscopy and SEC/MALLS in comparison with hydroxyethyl starch
- The properties of starch derivs. which may be used as plasma AB substitutes, are dependent upon the mol. structure. Seven acetyl starch (AS) samples were determined and compared with results from hydroxyethyl starch (HES) samples. The molar masses and their distributions were determined with the combination of size exclusion chromatog. and light scattering. Slightly asym. distributions were determined with a polydispersity Mw /Mn .simeq. 2.4 and weight-average molar masses of Mw = 250,000-300,000g/mol for 6 AS samples and Mw/Mn .simeq. 3.6 and a weight-average molar mass of 766,000 g/mol for one AS sample. The average degrees of substitution (DS) and the substitution pattern were determined by high resolution MNR spectroscopy. The AS samples investigated had a DS of 0.42-0.81, comparable to HES, but the regionelective substitution pattern revealed differences. While for HES the position C-2 is preferred and the position C-3 has nearly no substituent, for AS both positions, C-2 and C-3, are substituted likewise. Degradability by  $\alpha$ -amylase was tested in the laboratory for As as well as for HES having nearly the same degree of substitution and molar mass, but C-2/C-6 = 2 for AS and C-2/C-6 = 21.4 for HES. An exponential decrease in the molar mass was observed over time, down to a limiting molar mass Mw .simeq. 50,000 g/mol for AS and Mw .simeq. 30,000 g/mol for HES, the degradation of AS occurred more slowly.
- AN 1998:771673 HCAPLUS <<LOGINID::20101007>>
- DN 130:29151
- TI Characterization of acetyl starch by means of NMR spectroscopy and SEC/MALLS in comparison with hydroxyethyl starch
- AU Heins, Dorothee; Kulicke, Werner-Michael; Kaeuper, Peter; Thielking, Heiko
- CS Institut Technische Makromolekulare Chemie, Universitaet Hamburg, Hamburg, D-20146, Germany
- SO Starch/Staerke (1998), 50(10), 431-437 CODEN: STARDD; ISSN: 0038-9056
- PB Wiley-VCH Verlag GmbH
- DT Journal
- LA English
- OSC.G 20 THERE ARE 20 CAPLUS RECORDS THAT CITE THIS RECORD (20 CITINGS)
- RE.CNT 20 THERE ARE 20 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L13 ANSWER 33 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI No coagulation disorders under high-dose volume therapy with low-molecular-weight hydroxyethyl starch
- Hydroxyethyl starch (HES) is often used for volume therapy. Since bleeding AB complications were reported repeatedly, a strict dose limitation of a maximum of 1500 mL 6% solution/day is recommended. Many indications require higher dosages. Bleeding complications are known to be caused by an acquired von Willebrand syndrome. It was shown that the accumulation of large mols. and their impairment in the coagulation system can be avoided by HES prepns. with a low in vivo mol. weight The effects of a high-dose therapy were not studied yet. The authors have investigated, how a 4-day high-dose therapy, using 3,000 mL 6% HES 70/0.5 on the 1st day and 1,500 mL on days 2-4, affects the coagulation system and hemorheol. parameters of acute stroke patients. Thromboplastin time, activated partial thromboplastin time and thrombin time showed no changes, except for a slight, clin. irrelevant change due to dilution The subunits of von Willebrand factor VIII showed no change. Hematocrit decreased from 42.3 to 37.4 after day 1, reaching 35.3% at the end of the therapy, demonstrating a substantial volume effect. Plasma viscosity and erythrocyte aggregation decreased slightly, however not significantly. Our study shows that even a high-dose therapy with 6% HES 70/0.5 has no influence on the coagulation system.
- AN 1998:430987 HCAPLUS <<LOGINID::20101007>>
- DN 129:62698
- OREF 129:12841a,12844a
- TI No coagulation disorders under high-dose volume therapy with low-molecular-weight hydroxyethyl starch
- AU Stoll, Martin; Treib, Johannes; Schenk, Joachim F.; Windisch, Florian; Haass, Anton; Wenzel, Ernst; Schimrigk, Klaus
- CS Dep. Neurology, Univ. Saarland, Homburg/Saar, D-66421, Germany
- SO Haemostasis (1998), Volume Date 1997, 27(5), 251-258 CODEN: HMTSB7; ISSN: 0301-0147
- PB S. Karger AG
- DT Journal
- LA English
- OSC.G 5 THERE ARE 5 CAPLUS RECORDS THAT CITE THIS RECORD (5 CITINGS)
- L13 ANSWER 34 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Polymeric compositions for purifying apolipoproteins
- AΒ The present invention relates to a composition for use in purification of apolipoprotein A (ApoA) or apolipoprotein E (ApoE), the composition comprising a first and a second polymeric material, wherein the first and second polymeric materials are immiscible in the primary aqueous solution, and wherein the second polymeric material is amphiphilic and water soluble. The resulting primary aqueous solution is maintained for a period of time sufficient for essentially separating the phases formed, and removing the phase containing the second polymeric material and the main portion of ApoA or ApoE. Subsequently, the second polymeric material is separated from ApoA or ApoE. The ApoA or ApoE obtained can be used for the manufacture of a medicament in the treatment of atherosclerosis and cardiovascular diseases, sepsis or peripheral atherosclerosis as well as in a method for treatment of atherosclerosis and cardiovascular diseases, sepsis or peripheral atherosclerosis when administered in a therapeutically effective amount The effect on purification and yield of primary aqueous 2-phase separation followed by

temperature-induced phase separation was studied by using ApoA-IM as the target protein. After cell removal, an  $E.\ coli$  fermentation solution containing Apo A-IM was

added to an aqueous solution containing 17% Reppal PES-100, 12% PEG-PPG copolymer and

3.5M urea. The degree of purification and the yield of Apo A-IM were calculated based on the ELISA results. 1998:183946 HCAPLUS <<LOGINID::20101007>> ΑN 128:261925 DN OREF 128:51763a,51766a TIPolymeric compositions for purifying apolipoproteins ΙN Ageland, Hans; Nystrom, Lena; Persson, Josefine; Tjerneld, Folke PΑ Pharmacia & Upjohn AB, Swed. SO PCT Int. Appl., 33 pp. CODEN: PIXXD2 DT Patent LA English FAN.CNT 1 APPLICATION NO. PATENT NO. KIND DATE \_\_\_\_ \_\_\_\_\_\_ A1 19980319 WO 1997-SE1501 WO 9811140 19970908 <--РΤ W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TR, TT, UA, UG, US, UZ, VN RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG US 6559284 В1 20030506 US 1997-924994 19970905 <--CA 2277199 Α1 19980319 CA 1997-2277199 19970908 <--AU 9741429 19980402 AU 1997-41429 19970908 <--Α 19990922 EP 942935 A1 EP 1997-939314 19970908 <--R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI JP 2000513945 JP 1998-513556 Т 20001024 19970908 <--19960911 <--PRAI SE 1996-3303 Α Р 19960926 <--US 1996-26740P W WO 1997-SE1501 19970908 <--ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT OSC.G 4 THERE ARE 4 CAPLUS RECORDS THAT CITE THIS RECORD (5 CITINGS) RE.CNT 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT L13 ANSWER 35 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN TΙ Polymeric compositions for purifying apolipoproteins AB The present invention relates to a process for purifying a hydrophobic or amphiphilic compound, by first mixing a starting material containing the hydrophobic or amphiphilic compound with a first polymeric material, water and at 1 of a second polymeric material and a surfactant, wherein the first polymeric material and the second polymeric material and/or surfactant are immiscible in the resulting primary aqueous solution The process further comprises maintaining the primary aqueous solution for a period of time sufficient for essentially separating the phases formed, and then removing the phase containing the main portion of the hydrophobic or amphiphilic compound and the second polymeric material and/or surfactant. The second polymeric material and/or surfactant are separated from the hydrophobic or amphiphilic compound, and subsequently recycled to the initial mixing step. The present invention further relates to a composition for use in purification of apolipoprotein

A (ApoA) or apolipoprotein E (ApoE), said composition comprising a first polymeric material and a surfactant, said first polymeric material and surfactant being immiscible in the primary aqueous solution obtained after

mixing

with water. ApoA or ApoE produced by this process can be used for the manufacture of a medicament in the treatment of atherosclerosis and cardiovascular diseases, sepsis or peripheral atherosclerosis as well as in a method for treatment of atherosclerosis and cardiovascular diseases, sepsis or peripheral atherosclerosis when administered in a therapeutically effective amount. The effect on purification and yield of

aqueous 2-phase separation followed by temperature-induced phase separation was studied by

using ApoA-IM as the target protein. After cell removal, an E. coli fermentation solution containing Apo A-IM was added to an aqueous solution containing 8% Reppal

PES-100, 16% Breox PAG-50A-1000 and 0-40% Triton X-100. The degree of purification and the yield of Apo A-IM after separation from Reppal PES-100 in the

primary step, and from Breox PAG-50A-1000 and Triton X-100 in the temperature-induced phase separation were determined by gel scanning with densitometer.

AN 1998:183934 HCAPLUS <<LOGINID::20101007>>

DN 128:261924

OREF 128:51763a,51766a

- TI Polymeric compositions for purifying apolipoproteins
- IN Ageland, Hans; Nystrom, Lena; Persson, Josefine; Tjerneld, Folke
- PA Pharmacia & Upjohn AB, Swed.
- SO PCT Int. Appl., 46 pp. CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

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US 2000-571683 A1 20000516 <--

ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT

- OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)
- RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L13 ANSWER 36 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Coaqulation disorders caused by hydroxyethyl starch
- AΒ A review with 86 refs. is given on coagulation disorders caused by hydroxyethyl starch (HES). Initially, HES was only characterized by its in vitro mol. weight (MW). This is not sufficient because HES is degraded in vivo. One relevant parameter that predicts the rate of enzymic breakdown is the degree of substitution, a measure of the average number of hydroxyethyl groups per Glc unit. The higher this degree of substitution, the slower the break-down. In addition, because the Glc units can be substituted at C 2, 3, and 6, different substitution patterns are possible. They are classified by their C2/C6 hydroxyethylation ratio. A higher C2/C6 ratio results in less metabolism of the starch in vivo and results in a larger in vivo MW. This in turn affects therapy, because the larger the in vivo MW, the longer is the duration of the volume effect of HES. Of particular importance is the fact that HES with a high in vivo MW affects factor VIII/von Willebrand factor which can lead to an acquired von Willebrand syndrome. During a 10-day volume therapy with a medium-MW HES 200, a form that is difficult to metabolize, we observed an 80% drop in factor VIII/von Willebrand factor. Therapy with a medium- ${\tt MW}$  HES 200, a form that is easily degraded, and therapy with a low-MW HES 70 did not result in a relevant decline of factor VIII/von Willebrand factor. This explains why hemorrhagic complications were observed repeatedly in the United States after therapy with HES infusions, some of them lethal. In the United States high-MW HES 480 which is difficult to degrade is most frequently used and results in a larger in vivo MW and subsequent decrease in factor VIII/von Willebrand factor levels. In Europe, medium-MW HES 200 that is easily degraded and low-MW HES 70 are preferred. In the future, HES should be characterized by the in vivo, not the in vitro
- AN 1997:603004 HCAPLUS <<LOGINID::20101007>>
- DN 127:242743
- OREF 127:47199a,47202a
- TI Coaquiation disorders caused by hydroxyethyl starch
- AU Treib, Johannes; Haass, Anton; Pindur, Gerhard
- CS Department Neurology, University Saarland, Homburg, D-66421, Germany
- SO Thrombosis and Haemostasis (1997), 78(3), 974-983 CODEN: THHADQ; ISSN: 0340-6245
- PB Schattauer
- DT Journal; General Review
- LA English
- OSC.G 42 THERE ARE 42 CAPLUS RECORDS THAT CITE THIS RECORD (42 CITINGS)
- L13 ANSWER 37 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Increased hemorrhagic risk after repeated infusion of highly substituted medium molecular weight hydroxyethyl starch
- AB Infusion of large vols. of high mol. weight hydroxyethyl starch (HES) has been known to lead to coagulation disorders. Medium mol. weight starch is considered a safe alternative, even after repeated administration. In 10 patients with cerebrovascular diseases, a 10-day hemodilution was carried out using 10% HES 200/ 0.62. Initially, a loading dose of 500 mL was administered once over 4560 min, followed by 500 mL maintenance dose per day for 10 days. Its high intravascular mol. weight (120,000 D) showed that cleavage of the starch is slowed due to the higher degree of

substitution. The continuous increase of HES-serum concentration to 27.7 mg/mL gave evidence of a cumulation of poorly degradable mols. Although this caused a prolonged volume effect, plasma viscosity and erythrocyte aggregation were influenced in an unfavorable way. The neg. effects were most evident in their influence on the coagulation system. Under therapy, a significant 42.8% increase in activated partial thromboplastin time occurred. Factor VIII:C, von Willebrand ristocetin cofactor and von Willebrand factor antigen dropped during the therapy below the hemostasiol. limit of 30%, and in some patients below 10%. A high degree of substitution, particularly after repeated infusion, leads to a cumulation of large mols. that are difficult to break down and which unfavorably affect rheol. and hemostasiol. parameters.

- AN 1997:93454 HCAPLUS <<LOGINID::20101007>>
- DN 126:246632
- OREF 126:47554h,47555a
- TI Increased hemorrhagic risk after repeated infusion of highly substituted medium molecular weight hydroxyethyl starch
- AU Treib, Johannes; Haass, Anton; Pindur, Gerhard; Grauer, Markus T.; Jung, Friedel; Wenzel, Ernst; Schimrigk, Klaus
- CS Department Neurology, University Saarland, Homburg, D-66421, Germany
- SO Arzneimittel-Forschung (1997), 47(1), 18-22 CODEN: ARZNAD; ISSN: 0004-4172
- PB Cantor
- DT Journal
- LA English
- OSC.G 11 THERE ARE 11 CAPLUS RECORDS THAT CITE THIS RECORD (11 CITINGS)
- L13 ANSWER 38 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI The influence of volume therapy and pentoxifylline infusion on circulating adhesion molecules in trauma patients
- AΒ Adhesion mols. appear to play a pivotal role in tissue damage secondary to the inflammatory process. Besides neutrophil-and endothelial-bound adhesion mols., soluble forms have been detected in the circulating blood. They seem to be good markers of endothelial damage, but they may also have other biol. functions. Plasma concns. of soluble adhesion mols. (endothelial leukocyte adhesion mols. (sELAM-1)), intercellular adhesion mol.-1 (sICAM-1), vascular cell adhesion mol.-1 (sVCAM-1), and granule membrane protein 140 (sGMP-140) were serially measured over 5 days by enzyme-linked immunosorbent assays (ELISA) in 45 consecutive trauma patients. These received, by random allocation, only either hydroxyethylstarch solution 10% (mean mol. weight 200 000 Da) (n = 15) or human albumin 20% (n = 15) for volume therapy. Another 15 patients without defined volume therapy received pentoxifylline continuously (1.2 mg.kg-1.h-1). Measurements were carried out on the day of admission to the intensive care unit (baseline) and during the next 5 days. At baseline, plasma concns. of all adhesion mols. were similar in all groups. In the hydroxyethyl starch group, sELAM-1 and sICAM-1 concns. decreased significantly reaching normal values during the study period whereas the mean (SD) values increased in the pentoxifylline group (sELAM-1: 71.1 (16.7) to 91.6 (17.8) ng.ml-1) and the albumin group (sICAM-1: 400 (81) to 749 (101) nq.ml-1). SVCAM-1 increased outside the normal range only in the human albumin group (to 760 ng.ml-1). SGMP-140 plasma concentration increased only in those receiving albumin (432 (85) to 550 (93) ng.ml-1) and this was significantly different to the other groups. None of the other hemodynamic or laboratory factors could be correlated

with plasma concns. of the adhesion mols. The authors conclude that volume therapy with hydroxyethyl starch resulted in a decrease in circulating adhesion mols. in the authors' trauma patients. In contrast, volume therapy with albumin did not exert this effect. Continuous infusion

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of pentoxifylline did not have a beneficial modulating action on
     circulating adhesion mols.
     1997:48369 HCAPLUS <<LOGINID::20101007>>
ΑN
    126:139669
DN
OREF 126:26839a,26842a
    The influence of volume therapy and pentoxifylline infusion on circulating
     adhesion molecules in trauma patients
ΑU
     Boldt, J.; Heesen, M.; Padberg, W.; Martin, K.; Hempelmann, G.
CS
     Department Anaesthesiology and Intensive Care Medicine,
     Justus-Liebig-University Giessen, Giessen, Germany
SO
     Anaesthesia (1996), 51(6), 529-535
     CODEN: ANASAB; ISSN: 0003-2409
PΒ
     Saunders
DT
     Journal
    English
LA
OSC.G
       8
              THERE ARE 8 CAPLUS RECORDS THAT CITE THIS RECORD (8 CITINGS)
RE.CNT 37
              THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS RECORD
             ALL CITATIONS AVAILABLE IN THE RE FORMAT
    ANSWER 39 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
L13
     Influence of low molecular weight hydroxyethyl starch
ТΤ
     (HES 40/0.5-0.55) on hemostasis and hemorheology
AΒ
     The rheol. parameters erythrocyte aggregation and plasma
     viscosity were lowered in hemodilution therapy with low mol. weight HES
     (56-61 kD) compared to therapy with high or medium mol. weight HES. HES does
     not effect hemostasis (thromboplastin time, thrombin time, fibrinogen
     concentration).
ΑN
     1996:659040 HCAPLUS <<LOGINID::20101007>>
DN
     126:42492
OREF 126:8241a,8244a
     Influence of low molecular weight hydroxyethyl starch
     (HES 40/0.5-0.55) on hemostasis and hemorheology
     Treib, Johannes; Haass, Anton; Pindur, Gerhard; Grauer, Markus T.;
ΑU
     Seyfert, Ulrich T.; Treib, Wolfgang; Wenzel, Ernst; Schimrigk, Klaus
CS
     Department Neurology, University Saarland, Homburg, D-66421, Germany
SO
     Haemostasis (1996), 26(5), 258-265
     CODEN: HMTSB7; ISSN: 0301-0147
РΒ
     Karger
DT
     Journal
LA
     English
OSC.G
       10
             THERE ARE 10 CAPLUS RECORDS THAT CITE THIS RECORD (10 CITINGS)
L13 ANSWER 40 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
TΙ
     All medium starches are not the same: influence of the degree of
     hydroxyethyl substitution of hydroxyethyl starch on plasma
     volume, hemorheologic conditions, and coagulation
AΒ
     After the administration of high vols. of high-mol.-weight starch
     (hetastarch), bleeding complications have repeatedly been observed Later
     studies showed that the application of medium-mol.-weight starch led to far
     fewer disturbances of the blood coagulation system. However, the
     relationships among the degree of hydroxyethyl substitution, the rate of
     degradation, and the average in vivo mol. weight have not been investigated.
     10-day hemodilution treatment (n = 20) was carried out using two
     medium-mol.-weight hydroxyethyl starches (HES) with a degree of hydroxyethyl
     substitution of 0.5 and 0.62, resp. (10% HES 200 was used for a
     substitution of 0.5 and 6% HES 200 for a substitution of 0.62). After a
     loading dose of 500 mL was administered, 1000 mL of HES was infused daily
     for 4 days, and then 500 mL was infused daily for 6 days. The more highly
     substituted starch was broken down more slowly and eliminated renally.
     This resulted in a higher intravascular mol. weight than for the less highly
     substituted HES (120 vs. 84 kDa) and a greater increase in serum
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concentration (20.3 vs. 9.0 mg/mL). Initially, the more highly substituted 6-percent HES had a lesser effect on plasma volume (p<0.01). Because of HES accumulation, there was no longer a significant difference between the starches by the end of treatment, even though a higher dose of the 10-percent low-substitution starch was infused. Six-percent HES caused an increase in plasma viscosity (+9%, p<0.01) that was due to an accumulation of macromols. Ten-percent HES 200/0.5 had no effect on the coaquiation system beyond the dilution effect. Six-percent HES, on the other hand, led to an acquired von Willebrand syndrome during the course of the 10-day therapy. Factor VIII function was reduced by 72.2 percent, von Willebrand ristocetin cofactor by 61.3 percent, and von Willebrand factor antigen by 64 percent (p<0.01). Thus, it is the intravascular and not the initial (in vitro) mol. weight that dets. the properties of HES. Especially after repeated administration, a high degree of hydroxyethyl substitution leads to an accumulation of macromols. that affect hemorheol. measures and the coagulation system just as adversely as high-mol.-weight starch does. Depending on the degree of substitution, medium-mol.-weight starches can have widely differing properties.

AN 1996:443362 HCAPLUS <<LOGINID::20101007>>

DN 125:158096

OREF 125:29307a,29310a

- TI All medium starches are not the same: influence of the degree of hydroxyethyl substitution of hydroxyethyl starch on plasma volume, hemorheologic conditions, and coagulation
- AU Treib, J.; Haass, A.; Pindur, G.; Grauer, M.T.; Wenzel, E.; Schimrigk, K.
- CS Department of Neurology, University of the Saarland, Homburg, Germany
- SO Transfusion (Bethesda, Maryland) (1996), 36(5), 450-455 CODEN: TRANAT; ISSN: 0041-1132
- PB American Association of Blood Banks
- DT Journal
- LA English
- OSC.G 39 THERE ARE 39 CAPLUS RECORDS THAT CITE THIS RECORD (39 CITINGS)
- L13 ANSWER 41 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- TI Influence of intravascular molecular weight of hydroxyethyl starch on platelets
- AΒ Complications concerning the blood coagulation have been observed repeatedly after administration of highly substituted, high mol. weight hydroxyethyl starch (HES), but it has not been examined as to how intravascular mol. weight and degree of substitution of HES influence platelet number and volume after repeated administration. Thirty patients with cerebrovascular diseases were treated for 10 days with hemo-dilution 500 To 1500 mL of HES 200/0.62 (n=10), HES 200/0.5 (n=10) or HES 40/0.5 (n=10) were infused daily. During the first days, the number of platelets was not lowered beyond the dilution effect, but at the end of the therapy the number of platelets had increased in all 3 groups beyond the initial value. Platelet volume was lowered significantly in the 3 groups. HES 200/0.62 caused the largest drop in platelet volume (-10%, p<0.01). A possible explanation could be that HES macromols. are attached to platelets or are phagocytized by them. The larger platelets are then broken down and, to compensate the loss, more thrombocytes are released. A correlation between the mol. weight of HES and the breakdown rate of the platelets can be suspected, because HES 200/0.62 had the highest intravascular mean mol. weight (121 kD) and the largest effect on platelet volume
- AN 1996:244160 HCAPLUS <<LOGINID::20101007>>
- DN 124:332408
- OREF 124:61377a,61380a
- TI Influence of intravascular molecular weight of hydroxyethyl starch on platelets
- AU Treib, J.; Haass, A.; Pindur, G.; Treib, W.; Wenzel, E.; Schimrigk, K.

- CS Dept. Neurology, University the Saarland, Homburg/Saar, D-66421, Germany
- European Journal of Haematology (1996), 56(3), 168-72 SO CODEN: EJHAEC; ISSN: 0902-4441
- РΒ Munksgaard
- DT Journal
- English LA
- OSC.G 21 THERE ARE 21 CAPLUS RECORDS THAT CITE THIS RECORD (21 CITINGS)
- L13 ANSWER 42 OF 42 HCAPLUS COPYRIGHT 2010 ACS on STN
- Effects of 6% hydroxyethyl starch and 3% modified fluid gelatin on
- intravascular volume and coagulation during intraoperative hemodilution AΒ In the perioperative period, artificial colloids are most often infused in doses of 500-1000 mL i.v. This randomized study compared the effects on intravascular volume and coagulation of .apprx.2000 mL of two isooncotic artificial colloids: 6% hydroxyethyl starch (HES; MW 200,000; substitution ratio 0.40-0.55) and 3% modified fluid gelatin (GEL). hypothesized more pronounced hypocoagulation with HES and a weaker intravascular volume effect of GEL. Forty-two patients, scheduled for primary total hip replacement, were allocated randomly to receive HES or GEL during acute normovolemic hemodilution and subsequent further intraoperative hemodilution. Blood samples were taken before and after 500 mL and 1000 mL of acute normovolemic hemodilution; intraoperatively after 20 mL/kg of artificial colloid and at the end of colloid infusion; on arrival in the recovery room; and 3 h later. We quantified: 1) coagulation variables; 2) blood loss; 3) hemodynamic stability; 4) necessary infusion volume; 5) interstitial extravasation, calculated from plasma vols. measured using albumin marked with technetium-99m and iodine-125, resp.; 6) percentage volume effect at the end of the study as well as hematocrit, total serum protein, and colloid osmotic pressure. Intraoperative volume therapy was guided by radial systolic pressure and systolic pressure variation, mixed venous Hb saturation in the pulmonary artery, and pulmonary capillary occlusion pressure. The following differences (HES vs GEL, P <0.05) were found: 382 vs 725 mL extravasation; 76% vs 56% intravascular volume expansion 7 h after the median point of artificial colloid infusion; 27% vs 29% hematocrit and 35 vs 45 g/L total serum protein on arrival in recovery; 4 vs 0 abnormal bleeding times (>900 s); 3437 vs 2778 mL blood loss. This study quantifies a poorer volume effect of GEL and a higher blood loss with HES. The higher blood loss was significant with one-tailed testing only. These observations warrant extra GEL infusion to avoid hemoreconcn. and caution with large dose HES.
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- TΙ Effects of 6% hydroxyethyl starch and 3% modified fluid gelatin on intravascular volume and coagulation during intraoperative hemodilution
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- University Hospitals, Katholieke Universiteit Leuven, Louvain, Belg. CS
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- РΒ Williams & Wilkins
- DT Journal
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- OSC.G THERE ARE 9 CAPLUS RECORDS THAT CITE THIS RECORD (9 CITINGS)